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(FILE 'HOME' ENTERED AT 07:33:28 ON 24 MAR 2005)
SET COST OFF

FILE 'HCAPLUS' ENTERED AT 07:34:02 ON 24 MAR 2005

L1 1 S (WO2000-IT309 OR IT99-RM465)/AP,PRN
E FAVA D/AU
E DJACZENKO/AU
L2 38 S E4,E5
E STRUMILLO/AU
L3 3 S E8,E12
E FAVA D/AU
L4 5 S E3-E5
L5 14 S 5 METHYL 2 1 METHYLETHYL CYCLOHEXANOL
L6 2 S L5 AND 1 ALPHA 2 BETA 5 ALPHA

FILE 'REGISTRY' ENTERED AT 07:40:50 ON 24 MAR 2005

L7 4 S 2216-51-5 OR 2623-23-6 OR 89-78-1 OR 1490-04-6
L8 435 S C10H20O/MF AND 46.150.1/RID AND 1/NR
L9 213 S L8 AND CYCLOHEXANOL
L10 29 S L9 AND 5 METHYL
L11 17 S L10 AND 2 1 METHYLETHYL
L12 15 S L11 NOT LABELED
L13 16 S L7,L12
L14 14 S L10 NOT L13
L15 15 S L13 NOT ACETATE
L16 3 S L15 AND 1 ALPHA AND 2 BETA AND 5 ALPHA
L17 12 S L15 NOT L16

FILE 'HCAPLUS' ENTERED AT 07:48:08 ON 24 MAR 2005

L18 5879 S L16
L19 10732 S MENTHOL
L20 12031 S L5,L6,L18,L19

FILE 'REGISTRY' ENTERED AT 07:50:33 ON 24 MAR 2005

L21 1 S 76-03-9
L22 1 S 69-72-7

FILE 'HCAPLUS' ENTERED AT 07:51:11 ON 24 MAR 2005

L23 7287 S L21
L24 12797 S (TRICHLOROACETIC OR TRICHLORO ACETIC OR TRI CHLOROACETIC OR T
L25 29 S L20 AND L23,L24
L26 24150 S L22
L27 60605 S 2() (HYDROXYBENZOIC OR HYDROXY BENZOIC) ()ACID OR 2() (HYDROXYBE
L28 523 S L20 AND L26,L27
L29 7 S L25 AND L28

FILE 'REGISTRY' ENTERED AT 07:53:54 ON 24 MAR 2005

L30 2 S MENTHOL/CN

FILE 'HCAPLUS' ENTERED AT 07:54:06 ON 24 MAR 2005

L31 6340 S L30 OR L17
L32 19 S L31 AND L23,L24
L33 325 S L31 AND L26,L27
L34 8 S L28,L33 AND L25,L32
L35 8 S L29,L34

FILE 'REGISTRY' ENTERED AT 07:54:44 ON 24 MAR 2005

L36 1 S 25322-68-3
L37 1 S 9002-89-5
L38 1 S 112-60-7
L39 1 S 2615-15-8

L40 4 S 124-94-7 OR 378-44-9 OR 83-43-2 OR 50-02-2
L41 1 S 64-17-5
L42 1 S 7647-14-5

FILE 'HCAPLUS' ENTERED AT 07:56:05 ON 24 MAR 2005

L43 7 S L35 AND L36-L42
L44 1 S L35 AND (PVA OR POLYVINYLALCOHOL OR POLYVINYL ALCOHOL OR POLY
L45 1 S L35 AND (PEG OR POLYETHYLENEGLYCOL OR POLYETHYLENEOXIDE OR PO
L46 2 S L35 AND (TETRAETHYLENEGLYCOL OR TETRAETHYLENE GLYCOL OR TETR
L47 1 S L35 AND (HEXAETHYLENEGLYCOL OR HEXAETHYLENE GLYCOL OR HEXA ET
L48 2 S L35 AND ?CORTICOSTER?
L49 1 S L35 AND (TRIAMCINOLON? OR BETAMETHASON? OR BETA METHASON? OR
L50 5 S L35 AND (ETOH OR ETHANOL OR ETHYLALCOHOL OR ETHYL ALCOHOL)
L51 3 S L35 AND (NACL OR (NA OR SODIUM) ()CHLORIDE)
L52 3 S L35,L43-L51 AND PHARMACEUT?/SC,SX,CW,BI
L53 1 S L2-L4 AND L20,L31
L54 3 S L1,L52,L53

FILE 'REGISTRY' ENTERED AT 08:01:55 ON 24 MAR 2005

L55 15 S L15,L17,L30
SEL RN
L56 355 S E1-E15/CRN
L57 0 S L56 AND 76-03-9/CRN
L58 4 S L56 AND 69-72-7/CRN

=> fil hcaplus

FILE 'HCAPLUS' ENTERED AT 08:03:57 ON 24 MAR 2005
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FILE COVERS 1907 - 24 Mar 2005 VOL 142 ISS 13
FILE LAST UPDATED: 23 Mar 2005 (20050323/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d l54 all hitstr tot

L54 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2005 ACS on STN
AN 2003:173431 HCAPLUS
DN 138:226721
ED Entered STN: 07 Mar 2003
TI **Pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation
IN Joshi, Yatindra; Somma, Russell
PA Novartis AG, Switz.; Novartis Pharma GmbH
SO PCT Int. Appl., 18 pp.
CODEN: PIXXD2
DT Patent
LA English

IC ICM A61K031-4458
 ICS A61K031-137; A61K047-00; A61K031-4458; A61K031-19
 CC 63-6 (Pharmaceuticals)
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2003018015	A1	20030306	WO 2002-EP9660	20020829
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LT, LU, LV, MA, MD, MK, MN, MX, NO, NZ, OM, PH, PL, PT, RO, RU, SE, SG, SI, SK, TJ, TM, TN, TR, TT, UA, US, UZ, VC, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR				
	US 2003049272	A1	20030313	US 2001-942809	20010830
	US 2003147975	A1	20030807	US 2003-385372	20030310
PRAI	US 2001-942809	A	20010830		

CLASS

	PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
	WO 2003018015	ICM	A61K031-4458
		ICS	A61K031-137; A61K047-00; A61K031-4458; A61K031-19
	US 2003049272	ECLA	A61K009/00M18B; A61K009/20H4; A61K009/48H4
	US 2003147975	ECLA	A61K009/00M18B; A61K009/20H4; A61K009/48H4
AB	A pharmaceutical composition which reduces or eliminates the drug abuse potential of central nervous system stimulant comprising: (a) a central nervous system stimulant selected from the group consisting of methylphenidate, amphetamine, methamphetamine, and combinations thereof; and (b) a mucous membrane irritant selected from the group consisting of organic and inorg. acid, salt, ketone, nitrite, sulfide, bisulfate, persulfate, glycerophosphate, hypophosphate, borate, titanate, amino acid, peptide, and combinations thereof, wherein the mucous membrane irritant produces irritation when contacted with the skin or mucous membrane. The present invention is based on the discovery that a central nervous system stimulant, such as methylphenidate, in combination with a mucous membrane irritant, such as citric acid, reduces or eliminates potential drug abuse by producing "irritation" when contacted with the dermis layer of skin or mucous membrane, and thus, prevents nasal absorption and/or injectability of the drug. Formulations of chewable tablets containing 2.5% methylphenidate and 10% citric acid is disclosed.		
ST	pharmaceutical drug abuse CNS stimulant mucous membrane irritation		
IT	Balsams		
	RL: ADV (Adverse effect, including toxicity); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (Peru; pharmaceutical composition for preventing drug abuse by producing mucous membrane irritation)		
IT	Drugs of abuse (abuse of; pharmaceutical composition for preventing drug abuse by producing mucous membrane irritation)		
IT	Quaternary ammonium compounds, biological studies		
	RL: ADV (Adverse effect, including toxicity); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (alkylbenzyltrimethyl, chlorides; pharmaceutical composition for preventing drug abuse by producing mucous membrane irritation)		
IT	Drug delivery systems (capsules; pharmaceutical composition for preventing drug abuse by producing mucous membrane irritation)		
IT	Drug delivery systems (emulsions; pharmaceutical composition for preventing drug abuse by producing mucous membrane irritation)		
IT	Drug delivery systems		

(granules; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Acids, biological studies
 RL: ADV (Adverse effect, including toxicity); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)
 (inorg.; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Acids, biological studies
 RL: ADV (Adverse effect, including toxicity); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)
 (organic; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Capsicum
 Irritants
 Mucous membrane
 Nervous system stimulants
 Podophyllum (plant)
 (**pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Amino acids, biological studies
 Carboxylic acids, biological studies
 Coal tar
 Ketones, biological studies
 Peptides, biological studies
 Salts, biological studies
 Titanates
 RL: ADV (Adverse effect, including toxicity); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)
 (**pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Tar
 RL: ADV (Adverse effect, including toxicity); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)
 (pine; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Drug delivery systems
 (powders; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Drug delivery systems
 (sachets; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Drug delivery systems
 (solids; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Drug delivery systems
 (solns.; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Balsams
 RL: ADV (Adverse effect, including toxicity); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)
 (storax; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Drug delivery systems
 (suspensions; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Drug delivery systems
 (tablets, chewable; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Drug delivery systems
 (tablets; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT Balsams
 RL: ADV (Adverse effect, including toxicity); BUU (Biological use,

unclassified); BIOL (Biological study); USES (Uses)

(tolu; **pharmaceutical** composition for preventing drug abuse by producing mucous membrane irritation)

IT 50-21-5, Lactic acid, biological studies 51-21-8, Fluorouracil
56-25-7, Cantharidin 57-03-4 **64-17-5, Ethanol**,
biological studies 64-19-7, Acetic acid, biological studies 67-63-0,
Isopropanol, biological studies **69-72-7, Salicylic**
acid, biological studies **76-03-9**,
Trichloroacetic acid, biological studies 76-22-2,
Camphor 77-92-9, Citric acid, biological studies 79-10-7, Propenoic
acid, biological studies 79-14-1, Glycolic acid, biological studies
89-05-4, Pyromellitic acid 89-32-7, Pyromellitic dianhydride
89-78-1, Menthol 90-64-2, Mandelic acid 90-80-2,
Gluconolactone 94-36-0, Benzoyl peroxide, biological studies 96-26-4,
Dihydroxyacetone 97-59-6, Allantoin 108-46-3, Resorcinol, biological
studies 108-95-2, Phenol, biological studies 110-16-7, Maleic acid,
biological studies 118-56-9, Homosalate 123-19-3, Dipropylketone
127-07-1, Hydroxyurea 127-17-3, Pyruvic acid, biological studies
131-53-3, Dioxybenzone 139-33-3, Ethylenediaminetetraacetic acid
disodium salt 144-55-8, Sodium bicarbonate, biological studies
302-79-4, Retinoic acid 518-28-5, Podofilox 526-83-0, Tartaric acid
526-95-4, Gluconic acid 1143-38-0, Anthralin 1310-58-3, Potassium
hydroxide, biological studies 1321-11-5, Aminobenzoic acid 5466-77-3,
Octyl methoxycinnamate 6915-15-7, Malic acid 7446-70-0, Aluminum
chloride, biological studies **7647-14-5, Sodium**
chloride, biological studies 7697-37-2, Nitric acid, biological
studies 7722-84-1, Hydrogen peroxide, biological studies 7761-88-8,
Silver nitrate, biological studies 8029-68-3, Ichthammol 11129-12-7,
Borate 14797-65-0, Nitrite, biological studies 14996-02-2, Bisulfate,
biological studies 15092-81-6, Peroxydisulfate ((SO3)2O22-)
16566-52-2, Hypophosphate 18496-25-8, Sulfide 56093-45-9, Selenium
sulfide 70424-62-3 85791-94-2 92348-62-4, Hydroxy octanoic acid
126094-21-1 201596-35-2, EP02 500717-39-5, 2,4,6,8-Nonatetraenoic acid
500718-56-9

RL: ADV (Adverse effect, including toxicity); BUU (Biological use,
unclassified); BIOL (Biological study); USES (Uses)

(**pharmaceutical** composition for preventing drug abuse by producing
mucous membrane irritation)

IT 113-45-1, Methylphenidate 300-62-9, Amphetamine 537-46-2,
Methamphetamine

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(**pharmaceutical** composition for preventing drug abuse by producing
mucous membrane irritation)

RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE

- (1) Alza Corp; WO 9962496 A 1999 HCAPLUS
- (2) Chungi, S; WO 0059479 A 2000 HCAPLUS
- (3) Dariani, M; US 5908850 A 1999 HCAPLUS
- (4) Mantelle, J; US 6210705 B1 2001 HCAPLUS
- (5) Pozuelo, J; US 4117161 A 1978 HCAPLUS

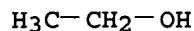
IT **64-17-5, Ethanol**, biological studies **69-72-7**,
Salicylic acid, biological studies **76-03-9**,
Trichloroacetic acid, biological studies **89-78-1**
, Menthol 7647-14-5, Sodium chloride
, biological studies

RL: ADV (Adverse effect, including toxicity); BUU (Biological use,
unclassified); BIOL (Biological study); USES (Uses)

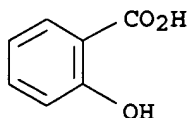
(**pharmaceutical** composition for preventing drug abuse by producing
mucous membrane irritation)

RN 64-17-5 HCAPLUS

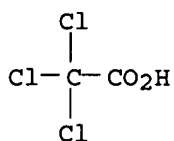
CN Ethanol (9CI) (CA INDEX NAME)



RN 69-72-7 HCAPLUS
 CN Benzoic acid, 2-hydroxy- (9CI) (CA INDEX NAME)

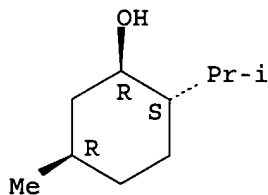


RN 76-03-9 HCAPLUS
 CN Acetic acid, trichloro- (8CI, 9CI) (CA INDEX NAME)



RN 89-78-1 HCAPLUS
 CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1R,2S,5R)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



RN 7647-14-5 HCAPLUS
 CN Sodium chloride (NaCl) (9CI) (CA INDEX NAME)

Cl-Na

L54 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2005 ACS on STN
 AN 2001:63814 HCAPLUS
 DN 134:120956
 ED Entered STN: 26 Jan 2001
 TI **Pharmaceutical** composition for topical application for skin injury treatment
 IN **Strumillo Djaczenko, Maria; Fava, Danila; Djaczenko, Wiktor**
 PA Italy
 SO PCT Int. Appl., 13 pp.
 CODEN: PIXXD2
 DT Patent
 LA English
 IC ICM A61K031-00
 CC 63-6 (Pharmaceuticals)
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2001005387	A2	20010125	WO 2000-IT309	20000721 <--
	WO 2001005387	A3	20010913		
	W:			AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM	
	RW:			GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG	
	IT 1306171	B1	20010530	IT 1999-RM465	19990721 <--
	CA 2379653	AA	20010125	CA 2000-2379653	20000721 <--
	EP 1218063	A2	20020703	EP 2000-951845	20000721 <--
	EP 1218063	B1	20040414		
	R:			AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL	
	AT 264125	E	20040415	AT 2000-951845	20000721 <--
PRAI	IT 1999-RM465	A	19990721	<--	
	WO 2000-IT309	W	20000721	<--	

CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
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WO 2001005387	ICM	A61K031-00
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AB **Pharmaceutical** compns. for topical use comprise a mixture, in a suitable polymer, of trichloroacetic acid, 2-hydroxybenzoic acid, menthol and, if desired, other **pharmaceutically** acceptable adjuvants and excipients and are used for the treatment of cutaneous injuries. A composition was prepared containing trichloroacetic acid in hexaethylene glycol, 2-hydroxybenzoic acid in hexaethylene glycol and menthol in ethanol.

ST topical **pharmaceutical** skin injury

IT Medical goods

(dressings; topical **pharmaceuticals** for skin injury treatment)

IT Skin, disease

(injury; topical **pharmaceuticals** for skin injury treatment)

IT Polyoxyalkylenes, biological studies

RL: MOA (Modifier or additive use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(topical **pharmaceuticals** for skin injury treatment)

IT **Corticosteroids**, biological studies

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(topical **pharmaceuticals** for skin injury treatment)

IT Drug delivery systems

(topical; topical **pharmaceuticals** for skin injury treatment)

IT 64-17-5, Ethanol, biological studies 112-60-7,

Tetraethylene glycol 2615-15-8,

Hexaethylene glycol 7647-14-5, Sodium

chloride, biological studies 9002-89-5,

Polyvinyl alcohol 25322-68-3, Peg

RL: MOA (Modifier or additive use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(topical **pharmaceuticals** for skin injury treatment)

IT 50-02-2, Dexamethasone 69-72-7, 2-

Hydroxybenzoic acid, biological studies 76-03-9

, Trichloroacetic acid, biological studies

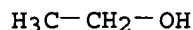
83-43-2, Methylprednisolone 89-78-1,

Menthol 124-94-7, Triamcinolone

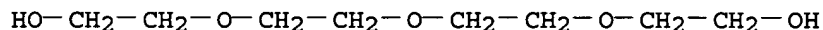
378-44-9, Betamethasone

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (topical **pharmaceuticals** for skin injury treatment)

IT 64-17-5, **Ethanol**, biological studies 112-60-7,
Tetraethylene glycol 2615-15-8,
Hexaethylene glycol 7647-14-5, **Sodium**
chloride, biological studies 9002-89-5,
Polyvinyl alcohol 25322-68-3, **Peg**
 RL: MOA (Modifier or additive use); THU (Therapeutic use); BIOL
 (Biological study); USES (Uses)
 (topical **pharmaceuticals** for skin injury treatment)
 RN 64-17-5 HCAPLUS
 CN Ethanol (9CI) (CA INDEX NAME)

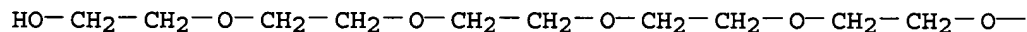


RN 112-60-7 HCAPLUS
 CN Ethanol, 2,2'-[oxybis(2,1-ethanediyl)oxy]bis- (9CI) (CA INDEX NAME)

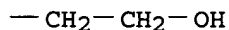


RN 2615-15-8 HCAPLUS
 CN 3,6,9,12,15-Pentaoxaheptadecane-1,17-diol (9CI) (CA INDEX NAME)

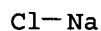
PAGE 1-A



PAGE 1-B



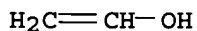
RN 7647-14-5 HCAPLUS
 CN Sodium chloride (NaCl) (9CI) (CA INDEX NAME)



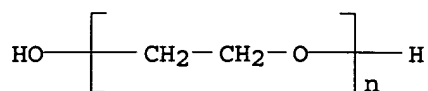
RN 9002-89-5 HCAPLUS
 CN Ethenol, homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 557-75-5
 CMF C2 H4 O

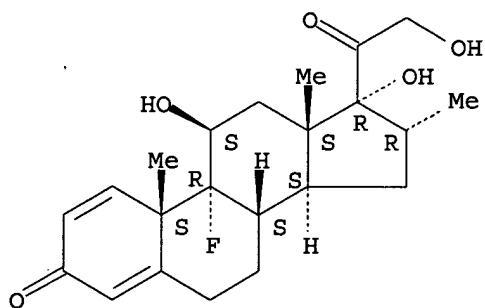


RN 25322-68-3 HCAPLUS
 CN Poly(oxy-1,2-ethanediyl), α -hydro- ω -hydroxy- (9CI) (CA INDEX NAME)

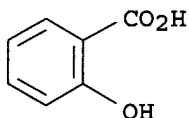


IT 50-02-2, Dexamethasone 69-72-7, 2-Hydroxybenzoic acid, biological studies 76-03-9, Trichloroacetic acid, biological studies 83-43-2, Methylprednisolone 89-78-1, Menthol 124-94-7, Triamcinolone 378-44-9, Betamethasone
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (topical **pharmaceuticals** for skin injury treatment)
 RN 50-02-2 HCAPLUS
 CN Pregna-1,4-diene-3,20-dione, 9-fluoro-11,17,21-trihydroxy-16-methyl-, (11 β ,16 α)- (9CI) (CA INDEX NAME)

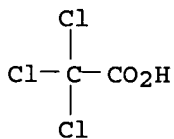
Absolute stereochemistry.



RN 69-72-7 HCAPLUS
 CN Benzoic acid, 2-hydroxy- (9CI) (CA INDEX NAME)

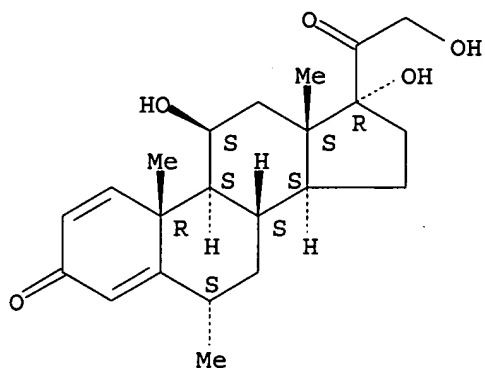


RN 76-03-9 HCAPLUS
 CN Acetic acid, trichloro- (8CI, 9CI) (CA INDEX NAME)



RN 83-43-2 HCAPLUS
 CN Pregna-1,4-diene-3,20-dione, 11,17,21-trihydroxy-6-methyl-, (6 α ,11 β)- (9CI) (CA INDEX NAME)

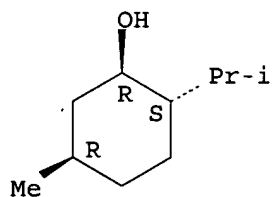
Absolute stereochemistry.



RN 89-78-1 HCAPLUS

CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1R,2S,5R)-rel- (9CI) (CA INDEX NAME)

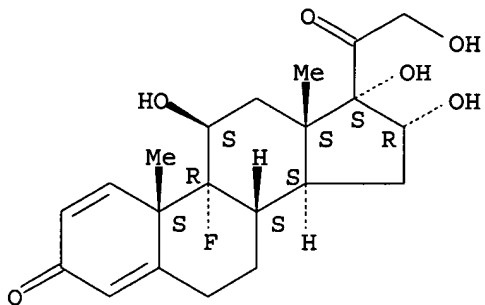
Relative stereochemistry.



RN 124-94-7 HCAPLUS

CN Pregna-1,4-diene-3,20-dione, 9-fluoro-11,16,17,21-tetrahydroxy-, (11 β ,16 α)- (9CI) (CA INDEX NAME)

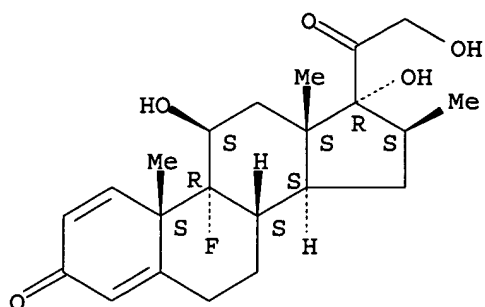
Absolute stereochemistry.



RN 378-44-9 HCAPLUS

CN Pregna-1,4-diene-3,20-dione, 9-fluoro-11,17,21-trihydroxy-16-methyl-, (11 β ,16 β)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L54 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 1998:799999 HCAPLUS

DN 130:43364

ED Entered STN: 22 Dec 1998

TI Pyridine thiols reverse mucocutaneous aging

IN Thornfeldt, Carl R.

PA Cellergy Pharmaceuticals Inc., USA

SO PCT Int. Appl., 21 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM A61K031-44

CC 63-6 (Pharmaceuticals)

Section cross-reference(s): 62

FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9853822	A1	19981203	WO 1998-US11270	19980602
	W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
	US 6071543	A	20000606	US 1998-89302	19980601
	AU 9877179	A1	19981230	AU 1998-77179	19980602
	US 6482839	B1	20021119	US 1998-145822	19980902
PRAI	US 1997-47360	P	19970602		
	US 1997-56282P	P	19970903		
	US 1998-89302	A	19980601		
	US 1997-47360P	P	19970602		
	US 1997-56290P	P	19970903		
	US 1997-58752P	P	19970912		
	WO 1998-US11270	W	19980602		

CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
WO 9853822	ICM	A61K031-44
WO 9853822	ECLA	A61K007/06C14B4; A61K007/48A2; A61K007/48C14D; A61K008/19; A61K008/20; A61K008/23; A61K008/26; A61K008/27; A61K008/29; A61K008/368; A61K008/49C6; A61K031/00; A61Q005/10; A61Q019/00; A61Q019/02; A61Q
US 6071543	ECLA	A61K007/06C14B4; A61K008/19; A61K008/20; A61K008/23; A61K008/26; A61K008/27; A61K008/28; A61K008/29; A61K008/68; A61K008/49C6; A61K031/00; A61Q005/10; A61Q019/00; A61Q019/02; A61Q019/08; A61K007/48A2;

US 6482839 ECLA A61K007/48C4D
 A61K007/06C14B4; A61K007/48A2; A61K007/48C14D;
 A61K008/19; A61K008/20; A61K008/23; A61K008/26;
 A61K008/27; A61K008/29; A61K008/368; A61K008/49C6;
 A61K031/00; A61Q005/10; A61Q019/00; A61Q019/02; A61Q

AB This invention provides compns. and methods for preventing and reversing the signs and symptoms of intrinsic and photo aging. The compns. include one or more pyridine-thiols and tautomers with attached metallic moieties. Administration of the compds. to aging skin and mucous membranes in topical formulations, either as the only active ingredient or in combination with other known active ingredients, prevents and reverses aging symptoms. Addnl. compns. for preventing and reversing aging contain one or more sulfides and oxides of these same metallic ions, either alone or in combination with other mols. known or suspected to exhibit age reversing properties. Topical formulations containing both a pyridine-thiol and tautomers with attached metallic moiety and a metallic sulfide and/or metallic oxide effectively prevent and reverse the signs and symptoms of mucocutaneous aging.

ST antiaging skin ointment pyridine thiol formulation

IT Solar radiation
 (aging induced by; pyridine thiols reverse mucocutaneous aging)

IT Skin, disease
 (aging, wrinkles; pyridine thiols reverse mucocutaneous aging)

IT Mucous membrane
 Skin, disease
 (aging; pyridine thiols reverse mucocutaneous aging)

IT Cosmetics
 Cosmetics
 (creams, wrinkle-preventing; pyridine thiols reverse mucocutaneous aging)

IT Carboxylic acids, biological studies
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
 (dicarboxylic; pyridine thiols reverse mucocutaneous aging)

IT Drug delivery systems
 (emulsions; pyridine thiols reverse mucocutaneous aging)

IT Drug delivery systems
 (gels; pyridine thiols reverse mucocutaneous aging)

IT Carboxylic acids, biological studies
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
 (hydroxy; pyridine thiols reverse mucocutaneous aging)

IT Skin
 (irregular pigmentation; pyridine thiols reverse mucocutaneous aging)

IT Drug delivery systems
 (lotions; pyridine thiols reverse mucocutaneous aging)

IT Anti-inflammatory agents
 (nonsteroidal; pyridine thiols reverse mucocutaneous aging)

IT Drug delivery systems
 (ointments, creams; pyridine thiols reverse mucocutaneous aging)

IT Drug delivery systems
 (ointments; pyridine thiols reverse mucocutaneous aging)

IT Carboxylic acids, biological studies
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
 (oxo; pyridine thiols reverse mucocutaneous aging)

IT Drug delivery systems
 (pastes; pyridine thiols reverse mucocutaneous aging)

IT Aging, animal
 Antioxidants

Cations
 Seborrhea
 Skin, neoplasm
 (pyridine thiols reverse mucocutaneous aging)
 IT Thiols (organic), biological studies
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
 (pyridine thiols reverse mucocutaneous aging)
 IT Aldehydes, biological studies
 Amides, biological studies
 Amino acids, biological studies
Corticosteroids, biological studies
 Esters, biological studies
 Flavanols
 Glucocorticoids
 Lactones
 Phenols, biological studies
 Salts, biological studies
 Sulfones
 Tocopherols
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
 (pyridine thiols reverse mucocutaneous aging)
 IT Drug delivery systems
 (solns.; pyridine thiols reverse mucocutaneous aging)
 IT Drug delivery systems
 (sprays; pyridine thiols reverse mucocutaneous aging)
 IT Drug delivery systems
 (suspensions; pyridine thiols reverse mucocutaneous aging)
 IT Drug delivery systems
 (topical; pyridine thiols reverse mucocutaneous aging)
 IT Acne
 (vulgaris; pyridine thiols reverse mucocutaneous aging)
 IT 110-86-1D, Pyridine, thiol derivs., biological studies
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
 (pyridine thiols reverse mucocutaneous aging)
 IT 50-21-5, biological studies 50-81-7, Ascorbic acid, biological studies
 53-86-1, Indomethacin 56-40-6, Glycine, biological studies 56-41-7, Alanine, biological studies 56-45-1, Serine, biological studies
 60-33-3, Linoleic acid, biological studies 64-18-6, Formic acid, biological studies 64-19-7, Acetic acid, biological studies 64-86-8, Colchicine 65-85-0, Benzoic acid, biological studies 68-26-8, Retinol 69-72-7, **Salicylic acid**, biological studies
 74-79-3, Arginine, biological studies 76-03-9, **Trichloroacetic acid**, biological studies 76-93-7, Benzilic acid, biological studies 77-92-9, Citric acid, biological studies 79-09-4, Propanoic acid, biological studies 79-11-8D, Chloroacetic acid, derivs. 79-14-1, Glycolic acid, biological studies 79-43-6, Dichloroacetic acid, biological studies 80-08-0, Dapsone 80-69-3, Tartronic acid 87-69-4, biological studies 87-73-0, Saccharic acid 89-83-8, Thymol 90-64-2, Mandelic acid 108-46-3, 1,3-Benzenediol, biological studies 108-95-2, Phenol, biological studies 110-15-6, Butanedioic acid, biological studies 110-17-8, 2-Butenedioic acid (2E)-, biological studies 116-31-4, Retinaldehyde 123-99-9, Azelaic acid, biological studies 127-17-3, Pyruvic acid, biological studies 136-77-6, Hexyl resorcinol 144-62-7, Ethanedioic acid, biological studies 300-85-6, 3-Hydroxybutyric acid 302-79-4, Tretinoin

463-40-1, Linolenic acid 470-82-6, Eucalyptol 473-81-4, Glyceric acid 501-30-4, Kojic acid 501-30-4D, Kojic acid, derivs. 526-99-8, Mucic acid 552-63-6, Tropic acid 685-73-4, Galacturonic acid 828-01-3 989-51-5, Epigallocatechin gallate 1198-69-2 1406-16-2D, Vitamin D, analogs **1490-04-6, Menthol** 2782-86-7, Heptonic acid 6556-12-3, Glucuronic acid 6915-15-7, Malic acid 7429-90-5, Aluminum, biological studies 7439-89-6, Iron, biological studies 7439-95-4, Magnesium, biological studies 7439-96-5, Manganese, biological studies 7440-02-0, Nickel, biological studies 7440-09-7, Potassium, biological studies 7440-20-2, Scandium, biological studies 7440-22-4, Silver, biological studies 7440-23-5, Sodium, biological studies 7440-24-6, Strontium, biological studies 7440-31-5, Tin, biological studies 7440-32-6, Titanium, biological studies 7440-38-2, Arsenic, biological studies 7440-43-9, Cadmium, biological studies 7440-47-3, Chromium, biological studies 7440-48-4, Cobalt, biological studies 7440-50-8, Copper, biological studies 7440-55-3, Gallium, biological studies 7440-56-4, Germanium, biological studies 7440-62-2, Vanadium, biological studies 7440-66-6, Zinc, biological studies 7440-67-7, Zirconium, biological studies 7440-70-2, Calcium, biological studies 7726-95-6, Bromine, biological studies 7782-49-2, Selenium, biological studies 13382-27-9, Galactonic acid 13463-41-7, Zinc pyrithione 13532-37-1, 4-Hydroxyvaleric acid 14915-37-8 15687-27-1, Ibuprofen 22071-15-4, Ketoprofen 29467-96-7, Pyridinethiol 34387-34-3 36322-90-4 56093-45-9, Selenium sulfide 62662-81-1, Methyl resorcinol 66664-10-6, Tetrahydroxypentanoic acid 74103-06-3, Ketorolac 216965-04-7
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (pyridine thiols reverse mucocutaneous aging)

RE.CNT 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

(1) Sakamoto, T; JP 61238716 A2 1986 HCAPLUS

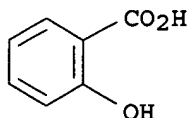
IT 69-72-7, **Salicylic acid**, biological studies

76-03-9, **Trichloroacetic acid**, biological studies **1490-04-6, Menthol**

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (pyridine thiols reverse mucocutaneous aging)

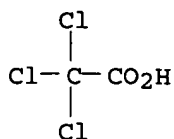
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CN Benzoic acid, 2-hydroxy- (9CI) (CA INDEX NAME)



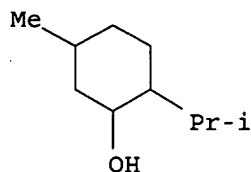
RN 76-03-9 HCAPLUS

CN Acetic acid, trichloro- (8CI, 9CI) (CA INDEX NAME)



RN 1490-04-6 HCAPLUS

CN Cyclohexanol, 5-methyl-2-(1-methylethyl)- (9CI) (CA INDEX NAME)



=> => fil reg

FILE 'REGISTRY' ENTERED AT 08:04:22 ON 24 MAR 2005
 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
 PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
 COPYRIGHT (C) 2005 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file
 provided by InfoChem.

STRUCTURE FILE UPDATES: 23 MAR 2005 HIGHEST RN 847137-45-5
 DICTIONARY FILE UPDATES: 23 MAR 2005 HIGHEST RN 847137-45-5

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 18, 2005

Please note that search-term pricing does apply when
 conducting SmartSELECT searches.

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*
* The CA roles and document type information have been removed from *
* the IDE default display format and the ED field has been added,   *
* effective March 20, 2005. A new display format, IDERL, is now    *
* available and contains the CA role and document type information. *
*
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Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more
 information enter HELP PROP at an arrow prompt in the file or refer
 to the file summary sheet on the web at:
<http://www.cas.org/ONLINE/DBSS/registryss.html>

=> => d ide can tot l61

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L61 ANSWER 1 OF 4 REGISTRY COPYRIGHT 2005 ACS on STN
RN 15356-60-2 REGISTRY
ED Entered STN: 16 Nov 1984
CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1S,2R,5S) - (9CI)
   (CA INDEX NAME)
OTHER CA INDEX NAMES:
CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, [1S-
   (1α,2β,5α)] -
CN Menthol, (1S,3S,4R) - (+) - (8CI)
OTHER NAMES:
CN (+)-Menthol
CN (1S,2R,5S) - (+) -Menthol
CN (1S,2R,5S) -Menthol
CN d-Menthol
FS STEREOSEARCH
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CI COM
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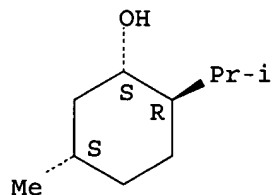
LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CAPLUS, CASREACT, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CSCHEM, HODOC*, HSDB*, IFICDB, IFIPAT, IFIUDB, IPA, NAPRALERT, TOXCENTER, USPAT2, USPATFULL

(*File contains numerically searchable property data)

Other Sources: DSL**, EINECS**, TSCA**

(**Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry. Rotation (+).



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

472 REFERENCES IN FILE CA (1907 TO DATE)

4 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

477 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 142:240047

REFERENCE 2: 142:197451

REFERENCE 3: 142:176415

REFERENCE 4: 142:134334

REFERENCE 5: 142:106156

REFERENCE 6: 142:93227

REFERENCE 7: 142:74760

REFERENCE 8: 142:32874

REFERENCE 9: 142:6900

REFERENCE 10: 142:6594

L61 ANSWER 2 OF 4 REGISTRY COPYRIGHT 2005 ACS on STN

RN 2216-51-5 REGISTRY

ED Entered STN: 16 Nov 1984

CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1R,2S,5R)- (9CI)
(CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, [1R-(1 α ,2 β ,5 α)]-

CN Menthol, (1R,3R,4S)-(-)- (8CI)

OTHER NAMES:

CN (-)-Menthol

CN (-)-Menthyl alcohol

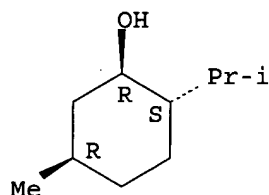
CN (-)-trans-p-Methan-cis-3-ol

CN (1R)-(-)-Menthol

CN (1R,2S,5R)-(-)-Menthol

CN (R)-(-)-Menthol
 CN 1R-Menthol
 CN 1-(-)-Menthol
 CN 1-Menthol
 CN Levomenthol
 CN NSC 62788
 FS STEREOSEARCH
 DR 98167-53-4
 MF C10 H20 O
 CI COM
 LC STN Files: ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*, BIOBUSINESS,
 BIOSIS, CA, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMINFORMRX,
 CHEMLIST, CIN, CSCHEM, DETHERM*, DIOGENES, DIPPR*, GMELIN*, HODOC*,
 HSDB*, IFICDB, IFIPAT, IFIUDB, IPA, MSDS-OHS, NAPRALERT, PROMT, PS,
 RTECS*, SPECINFO, TOXCENTER, USAN, USPAT2, USPATFULL
 (*File contains numerically searchable property data)
 Other Sources: DSL**, EINECS**, TSCA**
 (**Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry. Rotation (-).



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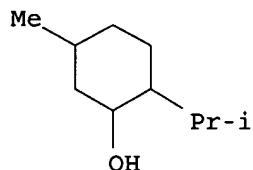
3042 REFERENCES IN FILE CA (1907 TO DATE)
 43 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
 3056 REFERENCES IN FILE CAPLUS (1907 TO DATE)

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 REFERENCE 3: 142:240047
 REFERENCE 4: 142:231851
 REFERENCE 5: 142:225699
 REFERENCE 6: 142:225255
 REFERENCE 7: 142:218968
 REFERENCE 8: 142:218727
 REFERENCE 9: 142:217709
 REFERENCE 10: 142:217661

L61 ANSWER 3 OF 4 REGISTRY COPYRIGHT 2005 ACS on STN
 RN 1490-04-6 REGISTRY
 ED Entered STN: 16 Nov 1984
 CN Cyclohexanol, 5-methyl-2-(1-methylethyl)- (9CI) (CA INDEX NAME)
 OTHER CA INDEX NAMES:
 CN Menthol (8CI)

OTHER NAMES:

CN 1-Methyl-4-isopropyl-3-cyclohexanol
CN 2-Isopropyl-5-methylcyclohexan-1-ol
CN 2-Isopropyl-5-methylcyclohexanol
CN 3-Hydroxy-p-menthane
CN 5-Methyl-2-(1-methylethyl)cyclohexanol
CN 5-Methyl-2-isopropylcyclohexanol
CN Menthyl alcohol
CN p-Menthan-3-ol
FS 3D CONCORD
MF C10 H20 O
CI COM
LC STN Files: ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
BIOTECHNO, CA, CABA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CEN,
CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSChem, CSNB, DDFU, DETHERM*,
DIOGENES, DRUGU, EMBASE, GMELIN*, HODOC*, IFICDB, IFIPAT, IFIUDB, IPA,
MEDLINE, MSDS-OHS, PDLCOM*, PHAR, PIRA, PROMT, RTECS*, SPECINFO,
TOXCENTER, USAN, USPAT2, USPATFULL, VETU, VTB
(*File contains numerically searchable property data)
Other Sources: DSL**, EINECS**, TSCA**
(**Enter CHEMLIST File for up-to-date regulatory information)



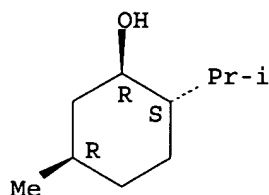
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3143 REFERENCES IN FILE CA (1907 TO DATE)
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1 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 142:246216
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REFERENCE 10: 142:225229

ED Entered STN: 16 Nov 1984
 CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1R,2S,5R)-rel- (9CI)
 (CA INDEX NAME)
 OTHER CA INDEX NAMES:
 CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-,
 (1 α ,2 β ,5 α)-
 CN Menthol, cis-1,3,trans-1,4- (8CI)
 CN Menthol, dl- (6CI)
 OTHER NAMES:
 CN (+)-Menthol
 CN (1R,2S,5R)-rel-5-Methyl-2-(1-methylethyl)cyclohexanol
 CN dl-Menthol
 CN Fisherman's Friend Lozenges
 CN Hexahydrothymol
 CN Menthacamphor
 CN Menthol
 CN Menthomenthol
 CN NSC 2603
 CN Peppermint camphor
 CN rac-Menthol
 CN Racementhol
 CN Therapeutic Mineral Ice
 CN Thymomenthol
 FS STEREOSEARCH
 DR 15356-70-4
 MF C10 H20 O
 CI COM
 LC STN Files: ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
 BIOTECHNO, CA, CAOLD, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS,
 CHEMINFORMRX, CHEMLIST, CIN, CSCHM, CSNB, DIOGENES, EMBASE, GMELIN*,
 HODOC*, HSDB*, IFICDB, IFIPAT, IFIUDB, MEDLINE, MRCK*, MSDS-OHS,
 NIOSHTIC, PDLCOM*, PIRA, PROMT, PS, RTECS*, SPECINFO, TOXCENTER, USAN,
 USPAT2, USPATFULL
 (*File contains numerically searchable property data)
 Other Sources: DSL**, EINECS**, TSCA**
 (**Enter CHEMLIST File for up-to-date regulatory information)

Relative stereochemistry.



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

2855 REFERENCES IN FILE CA (1907 TO DATE)
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 1 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

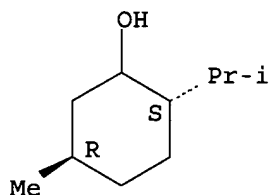
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REFERENCE 8: 142:213700
REFERENCE 9: 142:204735
REFERENCE 10: 142:204422

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L60 ANSWER 1 OF 11 REGISTRY COPYRIGHT 2005 ACS on STN
RN 477243-44-0 REGISTRY
ED Entered STN: 19 Dec 2002
CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (2R,5S)-rel- (9CI)
(CA INDEX NAME)
FS STEREOSEARCH
MF C10 H20 O
SR CA
LC STN Files: CA, CAPLUS, CASREACT, CHEMCATS, TOXCENTER

Relative stereochemistry.



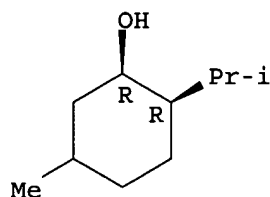
PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1907 TO DATE)
3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 141:139879
REFERENCE 2: 139:230691
REFERENCE 3: 137:310436

L60 ANSWER 2 OF 11 REGISTRY COPYRIGHT 2005 ACS on STN
RN 189076-46-8 REGISTRY
ED Entered STN: 16 May 1997
CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1R,2R)-rel- [partial]-
(9CI) (CA INDEX NAME)
FS STEREOSEARCH
MF C10 H20 O
SR CA
LC STN Files: CA, CAPLUS, CHEMCATS

Relative stereochemistry.



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 126:293451

L60 ANSWER 3 OF 11 REGISTRY COPYRIGHT 2005 ACS on STN

RN 64282-88-8 REGISTRY

ED Entered STN: 16 Nov 1984

CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1S,2S,5S)- (9CI)
(CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, [1S-(1 α ,2 α ,5 α)]-

OTHER NAMES:

CN (-)-Neoisomenthol

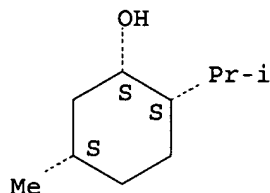
FS STEREOSEARCH

MF C10 H20 O

CI COM

LC STN Files: BEILSTEIN*, CA, CAPLUS, CASREACT, CHEMINFORMRX, CHEMLIST, IFICDB, IFIPAT, IFIUDB, SPECINFO, TOXCENTER, USPATFULL
(*File contains numerically searchable property data)

Absolute stereochemistry. Rotation (+).



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

19 REFERENCES IN FILE CA (1907 TO DATE)
19 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:217814

REFERENCE 2: 139:316174

REFERENCE 3: 138:39504

REFERENCE 4: 137:217182

REFERENCE 5: 137:41002

REFERENCE 6: 134:353480

REFERENCE 7: 129:39036
REFERENCE 8: 128:212406
REFERENCE 9: 124:305974
REFERENCE 10: 121:81222

L60 ANSWER 4 OF 11 REGISTRY COPYRIGHT 2005 ACS on STN

RN 23283-97-8 REGISTRY

ED Entered STN: 16 Nov 1984

CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1S,2R,5R)- (9CI)
(CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, [1S-(1 α ,2 β ,5 β)]-

CN Menthol, (1R,3S,4R)-(+)- (8CI)

OTHER NAMES:

CN (+)-Isomenthol

CN d-Isomenthol

FS STEREOSEARCH

MF C10 H20 O

CI COM

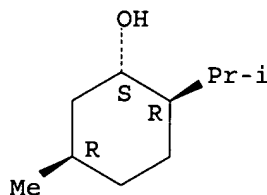
LC STN Files: AGRICOLA, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CAPLUS, CASREACT, CHEMCATS, CHEMINFORMRX, CHEMLIST, CSChem, HODOC*, IFICDB, IFIPAT, IFIUDB, SPECINFO, TOXCENTER, USPATFULL

(*File contains numerically searchable property data)

Other Sources: DSL**, EINECS**, TSCA**

(**Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry. Rotation (+).



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

108 REFERENCES IN FILE CA (1907 TO DATE)

108 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 142:218727
REFERENCE 2: 142:134146
REFERENCE 3: 141:260066
REFERENCE 4: 140:406589
REFERENCE 5: 140:217814
REFERENCE 6: 140:176684
REFERENCE 7: 140:12957
REFERENCE 8: 139:381216

REFERENCE 9: 139:350307

REFERENCE 10: 139:344933

L60 ANSWER 5 OF 11 REGISTRY COPYRIGHT 2005 ACS on STN

RN 20752-34-5 REGISTRY

ED Entered STN: 16 Nov 1984

CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1R,2R,5R)- (9CI)
(CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, [1R-(1 α ,2 α ,5 α)]-

CN Menthol, (1R,3R,4R)-(+)- (8CI)

OTHER NAMES:

CN (+)-Neoisomenthol

CN (R,R,R)-Menthol

FS STEREOSEARCH

MF C10 H20 O

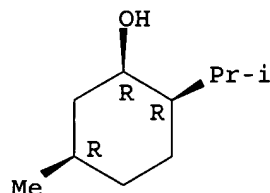
CI COM

LC STN Files: BEILSTEIN*, BIOBUSINESS, CA, CAPLUS, CASREACT, CHEMINFORMRX, CHEMLIST, HODOC*, IFICDB, IFIPAT, IFIUDB, TOXCENTER, USPAT2, USPATFULL
(*File contains numerically searchable property data)

Other Sources: DSL**

(**Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry. Rotation (-).



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

40 REFERENCES IN FILE CA (1907 TO DATE)

40 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 142:240063

REFERENCE 2: 141:411103

REFERENCE 3: 140:217814

REFERENCE 4: 139:316174

REFERENCE 5: 138:384980

REFERENCE 6: 137:41002

REFERENCE 7: 136:34778

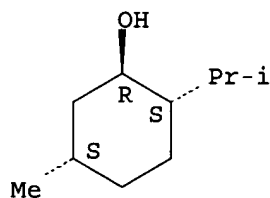
REFERENCE 8: 135:371876

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REFERENCE 10: 132:265305

L60 ANSWER 6 OF 11 REGISTRY COPYRIGHT 2005 ACS on STN
RN 20752-33-4 REGISTRY
ED Entered STN: 16 Nov 1984
CN **Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1R,2S,5S) - (9CI)**
(CA INDEX NAME)
OTHER CA INDEX NAMES:
CN **Cyclohexanol, 5-methyl-2-(1-methylethyl)-, [1R-(1 α ,2 β ,5 β)]-**
CN Menthol, (1S,3R,4S) - (8CI)
OTHER NAMES:
CN (-)-Isomenthol
FS STEREOSEARCH
MF C10 H20 O
CI COM
LC STN Files: BEILSTEIN*, CA, CAPLUS, CASREACT, CHEMINFORMRX, CHEMLIST, IFICDB, IFIPAT, IFIUDB, TOXCENTER, USPATFULL
(*File contains numerically searchable property data)

Absolute stereochemistry. Rotation (-).



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

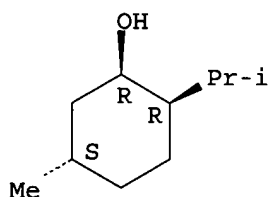
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REFERENCE 4: 139:344933
REFERENCE 5: 139:316174
REFERENCE 6: 138:384980
REFERENCE 7: 137:41002
REFERENCE 8: 136:216471
REFERENCE 9: 135:46313
REFERENCE 10: 132:265305

L60 ANSWER 7 OF 11 REGISTRY COPYRIGHT 2005 ACS on STN
RN 20747-49-3 REGISTRY
ED Entered STN: 16 Nov 1984
CN **Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1R,2R,5S) - (9CI)**
(CA INDEX NAME)
OTHER CA INDEX NAMES:
CN **Cyclohexanol, 5-methyl-2-(1-methylethyl)-, [1R-(1 α ,2 α ,5 β)]-**

CN Menthol, (1S,3R,4R) - (8CI)
 OTHER NAMES:
 CN (-) - (1R,2R,5S) -Neomenthol
 CN (-) -Neomenthol
 CN (1R,3R,4S) -p-Menth-3-ol
 CN 1-Neomenthol
 FS STEREOSEARCH
 MF C10 H20 O
 CI COM
 LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, CA, CAPLUS,
 CASREACT, CHEMCATS, CHEMINFORMRX, CHEMLIST, CSCHEM, HODOC*, IFICDB,
 IFIPAT, IFIUDB, TOXCENTER, USPATFULL
 (*File contains numerically searchable property data)

Absolute stereochemistry. Rotation (-).



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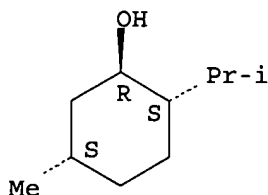
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 53 REFERENCES IN FILE CAPLUS (1907 TO DATE)

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 REFERENCE 4: 139:344933
 REFERENCE 5: 139:316174
 REFERENCE 6: 138:146579
 REFERENCE 7: 137:140102
 REFERENCE 8: 137:41002
 REFERENCE 9: 131:251896
 REFERENCE 10: 129:39036

L60 ANSWER 8 OF 11 REGISTRY COPYRIGHT 2005 ACS on STN
 RN 3623-52-7 REGISTRY
 ED Entered STN: 16 Nov 1984
 CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1R,2S,5S)-rel- (9CI)
 (CA INDEX NAME)
 OTHER CA INDEX NAMES:
 CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1 α ,2 β ,5 β)-
 (\pm)-
 CN Menthol, trans-1,3,cis-1,4-(\pm)- (8CI)
 OTHER NAMES:
 CN (\pm)-Isomenthol
 CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1 α ,2 β ,5 β)-

CN dl-Isomenthol
 CN Isomenthol
 FS STEREOSEARCH
 DR 490-99-3
 MF C10 H20 O
 LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA,
 CAOLD, CAPLUS, CASREACT, CHEMCATS, CHEMINFORMRX, CHEMLIST, CSCHEM, DDFU,
 DRUGU, EMBASE, GMELIN*, HODOC*, IFICDB, IFIPAT, IFIUDB, IPA, NAPRALERT,
 PS, RTECS*, SPECINFO, TOXCENTER, USPAT2, USPATFULL
 (*File contains numerically searchable property data)
 Other Sources: DSL**, EINECS**, TSCA**
 (**Enter CHEMLIST File for up-to-date regulatory information)

Relative stereochemistry.



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

346 REFERENCES IN FILE CA (1907 TO DATE)
 347 REFERENCES IN FILE CAPLUS (1907 TO DATE)
 1 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 142:182771
 REFERENCE 2: 142:151269
 REFERENCE 3: 142:37092
 REFERENCE 4: 141:354783
 REFERENCE 5: 141:162075
 REFERENCE 6: 141:156668
 REFERENCE 7: 141:106622
 REFERENCE 8: 141:690
 REFERENCE 9: 140:380241
 REFERENCE 10: 140:222991

L60 ANSWER 9 OF 11 REGISTRY COPYRIGHT 2005 ACS on STN

RN 3623-51-6 REGISTRY

ED Entered STN: 16 Nov 1984

CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1R,2R,5S)-rel- (9CI)
 (CA INDEX NAME)

OTHER CA INDEX NAMES:

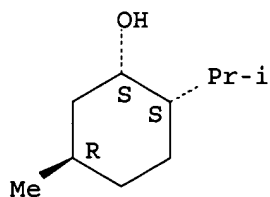
CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-,
 (1 α ,2 α ,5 β)-(±)-

CN Menthol, trans-1,3,trans-1,4-(±)- (8CI)

OTHER NAMES:

CN (±)-Neomenthol
CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-,
(1 α ,2 α ,5 β)-
CN dl-Neomenthol
CN neo-Menthol
CN Neomenthol
FS STEREOSEARCH
DR 491-01-0
MF C10 H20 O
CI COM
LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
BIOTECHNO, CA, CAOLD, CAPLUS, CASREACT, CHEMCATS, CHEMINFORMRX,
CHEMLIST, CSCHEM, DDFU, DRUGU, EMBASE, GMELIN*, HODOC*, IFICDB, IFIPAT,
IFIUDB, IPA, NAPRALERT, PS, RTECS*, SPECINFO, TOXCENTER, USPAT2,
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(*File contains numerically searchable property data)
Other Sources: DSL**, EINECS**, TSCA**
(**Enter CHEMLIST File for up-to-date regulatory information)

Relative stereochemistry.



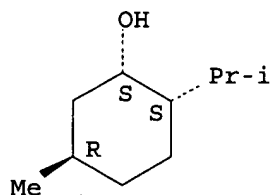
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472 REFERENCES IN FILE CA (1907 TO DATE)
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474 REFERENCES IN FILE CAPLUS (1907 TO DATE)
1 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 142:204422
REFERENCE 2: 142:182771
REFERENCE 3: 142:162597
REFERENCE 4: 142:113514
REFERENCE 5: 142:52174
REFERENCE 6: 142:37092
REFERENCE 7: 142:20054
REFERENCE 8: 142:6594
REFERENCE 9: 141:400453
REFERENCE 10: 141:257362

ED Entered STN: 16 Nov 1984
CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1S,2S,5R)- (9CI)
(CA INDEX NAME)
OTHER CA INDEX NAMES:
CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, [1S-(1 α ,2 α ,5 β)]-
CN Menthol, (1R,3S,4S)-(+)- (8CI)
OTHER NAMES:
CN (+)-neo-Menthol
CN (+)-Neomenthol
CN (1S,2S,5R)-Neomenthol
CN d-Neomenthol
FS STEREOSEARCH
MF C10 H20 O
CI COM
LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CAPLUS, CASREACT, CHEMCATS, CHEMINFORMRX, CHEMLIST, CSChem, GMELIN*, HODOC*, IFICDB, IFIPAT, IFIUDb, NAPRALERT, SPECINFO, TOXCENTER, USPAT2, USPATFULL
(*File contains numerically searchable property data)
Other Sources: DSL**, EINECS**, TSCA**
(**Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry. Rotation (+).



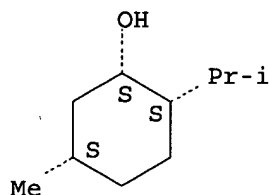
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REFERENCE 3: 141:314455
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REFERENCE 5: 140:249200
REFERENCE 6: 140:217814
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REFERENCE 8: 140:76900
REFERENCE 9: 140:12957
REFERENCE 10: 139:344933

RN 491-02-1 REGISTRY
 ED Entered STN: 16 Nov 1984
 CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-, (1R,2R,5R)-rel- (9CI)
 (CA INDEX NAME)
 OTHER CA INDEX NAMES:
 CN Cyclohexanol, 5-methyl-2-(1-methylethyl)-,
 (1 α ,2 α ,5 α)-
 CN Menthol, cis-1,3,cis-1,4- (8CI)
 OTHER NAMES:
 CN (+)-Neoisomenthol
 CN dl-Neoisomenthol
 CN Neoisomenthol
 FS STEREOSEARCH
 DR 3623-53-8
 MF C10 H20 O
 CI COM
 LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOSIS, CA, CAOLD, CAPLUS,
 CASREACT, CHEMINFORMRX, CHEMLIST, EMBASE, GMELIN*, HODOC*, IFICDB,
 IFIPAT, IFIUDB, NAPRALERT, SPECINFO, TOXCENTER, USPATFULL
 (*File contains numerically searchable property data)
 Other Sources: EINECS**
 (**Enter CHEMLIST File for up-to-date regulatory information)

Relative stereochemistry.



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

174 REFERENCES IN FILE CA (1907 TO DATE)
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 174 REFERENCES IN FILE CAPLUS (1907 TO DATE)
 1 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 142:133471
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 REFERENCE 3: 141:356044
 REFERENCE 4: 140:326575
 REFERENCE 5: 140:217823
 REFERENCE 6: 140:133364
 REFERENCE 7: 140:99245
 REFERENCE 8: 139:396063
 REFERENCE 9: 139:316174
 REFERENCE 10: 139:257988

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FILE 'WPIX' ENTERED AT 08:27:15 ON 24 MAR 2005

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FILE LAST UPDATED: 21 MAR 2005 <20050321/UP>

MOST RECENT DERWENT UPDATE: 200519 <200519/DW>

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FOR DETAILS. <<<

=> d all abeq tech abex

L101 ANSWER 1 OF 1 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN

AN 2001-147259 [15] WPIX

DNC C2001-043574

TI Pharmaceutical composition useful for treating cutaneous injuries, e.g.
resulting from burns, and dermatitis from animal sting, contains
**trichloroacetic acid, 2-hydroxybenzoic
acid and 5-methyl-2-(1-
methylethyl)cyclohexanol** in polymer.

DC A96 B05

IN DJACZENKO, W; FAVA, D; STRUMILLO, D M; STRUMILLO DJACZENKO, M

PA (DJAC-I) DJACZENKO W; (FAVA-I) FAVA D; (STRU-I) STRUMILLO M; (DJAC-I)
STRUMILLO DJACZENKO M

CYC 94

PI WO 2001005387 A2 20010125 (200115)* EN 13 A61K031-00

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CU CZ DE DK DM DZ
EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK
LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG
SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2000064688 A 20010205 (200128) A61K031-00

IT 1306171 B 20010530 (200229) A61K045-00

EP 1218063 A2 20020703 (200251) EN A61P017-02

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
RO SE SI

EP 1218063 B1 20040414 (200426) EN A61P017-02

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DE 60009933 E 20040519 (200434) A61P017-02

EP 1218063 B8 20040922 (200462) EN A61P017-02

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

ADT WO 2001005387 A2 WO 2000-IT309 20000721; AU 2000064688 A AU 2000-64688 20000721; IT 1306171 B IT 1999-RM465 19990721; EP 1218063 A2 EP 2000-951845 20000721, WO 2000-IT309 20000721; EP 1218063 B1 EP 2000-951845 20000721, WO 2000-IT309 20000721; DE 60009933 E DE 2000-00009933 20000721, EP 2000-951845 20000721, WO 2000-IT309 20000721; EP 1218063 B8 EP 2000-951845 20000721, WO 2000-IT309 20000721

FDT AU 2000064688 A Based on WO 2001005387; EP 1218063 A2 Based on WO 2001005387; EP 1218063 B1 Based on WO 2001005387; DE 60009933 E Based on EP 1218063, Based on WO 2001005387; EP 1218063 B8 Based on WO 2001005387

PRAI IT 1999-RM465 19990721

IC ICM A61K031-00; A61K045-00; A61P017-02
ICS A61K031-19; A61K031-57; A61P017-12

ICI A61K031-57, A61K031:045, A61K031:19

AB WO 200105387 A UPAB: 20010317

NOVELTY - A novel pharmaceutical composition for topical use comprises a mixture, in a suitable polymer, of **trichloroacetic acid**, **2-hydroxybenzoic acid**, (1 alpha, 2 beta, 5 alpha)-5-methyl-2-(1-methylethyl)cyclohexanol (I), and, if desired, other adjuvants and excipients.

ACTIVITY - Dermatological; tranquilizer; vulnerary; antiinflammatory. The composition was applied topically in liquid form on 10 subjects affected by first and second superficial degree burns, occurred in the previous 24 hours, which extended over a mean cutaneous surface of 25 cm² (varying from 4 to 200 cm²). The application of the composition caused in all the subjects a quick re-absorption of the liquid contained in the phlyctenas (within 1 hour after the first application), with reduction of the associated erythema and painful symptomatology. In all the subjects affected by first degree burns the restitutio ad integrum occurred 2 days after the injury event by a single application of the composition. In the subjects where phlyctenas were generated an immediate amelioration of the injury after the first application and restitutio ad integrum within 4/5 days occurred, during which period the composition was again applied to accelerate the exfoliation of the injured cutis. During the treatment local or systemic side-effects were not observed and the patients reported the application of the composition to be absolutely painless.

MECHANISM OF ACTION - None given.

USE - The compositions can be used for the treatment of cutaneous injuries, e.g. resulting from mechanical traumas or surgical operations, burns, dermatitis both from animal sting and animal or poisonous plant contact (claimed). They can also be used in the therapy and prevention of hypertrophic cicatrices and keloids (claimed). They can also be used in aesthetic medicine, e.g. as exfoliating agents (claimed).

ADVANTAGE - The compositions are able to reduce dramatically and, in many cases, to result in a complete disappearance of the hypertrophic cicatrices and keloids. The topical application is completely painless and does not result in any discomfort for the patient being treated.

Dwg.0/0

FS CPI

FA AB; DCN

MC CPI: A05-H03A; A10-E09B2; A12-V01; B01-B02; B04-C03B; B04-C03C; B05-A01B; B05-C07; B10-C03; B10-C04E; B10-E04A; B10-E04D; B14-C03; B14-J01B4; B14-N17A; B14-N17B; B14-N17C; B14-R01

TECH UPTX: 20010317

TECHNOLOGY FOCUS - POLYMERS - Preferred Material: The polymer may be polyethylene glycols, e.g. tetraethylene glycol or hexaethylene glycol, polyvinyl alcohol, or polyoxyethylene alcohol.

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Composition: The concentration of **trichloroacetic acid** is preferably 20-45 % w/v. The concentration of **2-hydroxybenzoic acid** is preferably 10-30 % w/v. The adjuvants and excipients are preferably ethanol or sodium chloride. Preparation: The composition may be

prepared by:

- (1) preparing, in an anhydrous atmosphere, the distinct (A) and (B) mixtures, respectively, of (A) **trichloroacetic acid** (40-90 %) in a suitable polymer, and (B) **2-hydroxybenzoic acid** (20-60 %) in a suitable polymer;
- (2) mixing, by adding small subsequent portions, the same volumes of the 2 mixtures to obtain the (A)+(B) mixture;
- (3) adding a volume of a (I) saturated solution in anhydrous ethanol equal to 2% of the volume of the (A)+(B) mixture;
- (4) adding of NaCl to obtain a final concentration of about 1.2% w/v;
- (5) keeping the obtained composition in a bottle filled with anhydrous air at 30 degrees C in reduced pressure conditions and leaving at ambient temperature for about 24 hours.

TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Composition: The compositions may also contain a corticosteroid, e.g. triamcinolone, betamethasone, methylprednisolone or dexamethasone.

ABEX UPTX: 20010317

ADMINISTRATION - The compositions can be used in the form of an ointment, gel, foam, liquid preparations, or medicated plaster. No dose given.

EXAMPLE - A mixture (A) containing **trichloroacetic acid** in hexaethylene glycol (apparent molecular weight (MW) 290) (50 % w/v) in anhydrous atmosphere (in a chemical hood in the presence of phosphorous pentoxide) was prepared, by adding the acid to the polymer portionwise and under continuous stirring to enhance the mixing. A mixture (B) containing **2-hydroxybenzoic acid** (as granules having diameter at most 50 microm) in hexaethylene glycol (apparent MW 290) (40 % w/v) was prepared. The acid was added to the polymer portionwise and under continuous stirring to enhance the mixing. After the preparation of the mixtures same volumes were mixed to obtain a (A)+(B) mixture (50 % v/v). The mixture was prepared using a 1000 ml tightly closed glass container on whose bottom (A) (10 ml) were poured. Then (B) (1 ml) and component (A), respectively, were stratified in the order. The amounts of (A) and (B) were again poured alternatively layer by layer to a 990 ml volume upon which the filling of the container was completed by pouring 10 ml of (B). A volume of a (1alpha, 2beta, 5alpha)-5-methyl-2-(1-methylethyl)cyclohexanol saturated solution in anhydrous ethanol equal to 2 % of volume of the (A)+(B) mixture was added; subsequently sodium chloride was added to obtain a final concentration of about 1.2 % w/v. Anhydrous air (through a solution of concentrated sulfuric acid) was blown within the thus obtained composition at 30 degrees C and reduced pressure, till air bubbles gurgled uniformly through the container, followed by its closing.

DEFINITIONS - An INDEPENDENT CLAIM is also included for the preparation of the composition.

=> => fil uspatful

FILE 'USPATFULL' ENTERED AT 08:42:19 ON 24 MAR 2005

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FILE COVERS 1971 TO PATENT PUBLICATION DATE: 22 Mar 2005 (20050322/PD)

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HIGHEST GRANTED PATENT NUMBER: US6871356

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CA INDEXING IS CURRENT THROUGH 22 Mar 2005 (20050322/UPCA)

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REVISED CLASS FIELDS (/NCL) LAST RELOADED: Feb 2005

USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Feb 2005

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>>> applications.  USPAT2 contains full text of the latest US      <<<
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>>> publications.  The publication number, patent kind code, and    <<<
>>> publication date for all the US publications for an invention  <<<
>>> are displayed in the PI (Patent Information) field of USPATFULL <<<
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>>> /PK, etc.                                                       <<<

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>>> enter this cluster.                                             <<<
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>>> classifications, or claims, that may potentially change from    <<<
>>> the earliest to the latest publication.                          <<<

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This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d l130 bib ab kwic tot

L130 ANSWER 1 OF 8 USPATFULL on STN

```

AN      2004:202914  USPATFULL
TI      Cosmetics or external preparations for skin
IN      Iwasaki, Keiji, Kanagawa, JAPAN
        Kitazawa, Manabu, Kanagawa, JAPAN
        Sakamoto, Kazutami, Kanagawa, JAPAN
PI      US 2004156802      A1  20040812
AI      US 2004-469985      A1  20040413 (10)
        WO 2002-JP2082      20020306
PRAI    JP 2001-70322      20010313
DT      Utility
FS      APPLICATION
LREP    OBLON, SPIVAK, MCCLELLAND, MAIER & NEUSTADT, P.C., 1940 DUKE STREET,
        ALEXANDRIA, VA, 22314
CLMN    Number of Claims: 11
ECL     Exemplary Claim: 1
DRWN    No Drawings
LN.CNT  1478
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB      According to the present invention, there is provided cosmetics or
        external preparations for skin which comprises, as the effective
        ingredient, a chemical peeling agent, a bactericide, an anionic
        surfactant or a cationic surfactant component in combination with a
        compound capable of mitigating the irritation, inflammation, etc. of the
        skin caused by these components.

```

The present invention is skin cosmetics, external preparations for skin or hair cosmetics which are characterized by containing the following components (A) and (B) or it is skin cosmetics or external preparations for skin which are characterized by containing the following components (C) and (D)

Components (A) and (C): one or more members selected from a cystine derivative and salt thereof.

Component (B): one or more of members selected from a chemical peeling agent, a bactericide and an anionic surfactant

Component (C): a cationic surfactant

AN 2004:202914 USPATFULL

SUMM . . . citric acid, isocitric acid, 2, 3, 4, 5-tetrahydroxyhexane-1,6-dioic acid (glucaric acid, mannaric acid, saccharic acid, mucic acid, etc.) quinic acid, **salicylic acid**, tropic acid, trethochoanic acid, citramalic acid, agaricic acid, aleuritic acid, pantoic acid, lactobiononic acid, hexuronic acid, etc. As other chemical. . . phenylpyruvic acid, methyl phenylpyruvate, ethyl phenylpyruvate, 2-ketobutyric acid, 2-ketopentanoic acid, 2-ketohexanoic acid, 2-ketoheptanoic acid, 2-ketooctanoic acid, 2-ketododecanoic acid, methyl 2-ketooctanoate; **trichloroacetic acid**, resorcinol, phenol and the like. Furthermore, there can be taken natural products or natural extracts containing the hydroxy acid or. . .

SUMM . . . phenol, chlorophenol, chloro-m-cresol, p-chloro-m-xylene, isopropyl methyl phenol, resorcinol, resorcinol acetate, o-phenylphenol, phenoxyethanol, thymol, cresol, hinokitiol, thioxolone, benzoic acid, sodium benzoate, **salicylic acid**, sodium salicylate, dehydroacetic acid and salts thereof, sorbic acid and salts thereof, hexachlorophene, trichlorohydroxydiphenyl ether (triclosan), trichlorocarbanilide, halocarboxylic acid, monoethanolamide. . .

SUMM . . . N-long chain acylalanine, N-long chain acylsarcosine, N-long chain acyl- β -alanine, N-long chain acyl-N-methyl- β -alanine and the like; ether carboxylates such as sodium **polyoxyethylene** (3E. O.) lauryl ether acetate, **polyoxyethylene** (3E. O.) lauryl ether acetic acid, sodium **polyoxyethylene** (3E. O.) tridecyl ether acetate, **polyoxyethylene** (3E. O.) tridecyl ether acetic acid, sodium **polyoxyethylene** (4.5E. O.) lauryl ether acetate, sodium **polyoxyethylene** (6E. O.) tridecyl ether acetate, **polyoxyethylene** (7E. O.) tridecyl ether acetic acid, sodium **polyoxyethylene** (3E. O.) stearyl ether acetate, sodium **polyoxyethylene** (3E. O.) octyl ether acetate, monoethanolamine **polyoxyethylene** (3E. O.) lauryl ether acetate and the like; alkyl sulfonates such as triethanolamine dodecylbenzenesulfonate, sodium dodecylbenzenesulfonate, sodium α -olefinsulfonate (carbon number. . . such as sulfosuccinic acid dioctyl ester sodium salt, sulfosuccinic acid lauryl ester disodium salt, sulfosuccinic acid tallow amide disodium salt, **polyoxyethylene** (1-SE. O.) sulfosuccinic acid lauryl ester disodium salt, **polyoxyethylene** (3E. O.) sulfosuccinic acid myristyl ester disodium salt, sulfosuccinic acid **polyoxyethylene** (5E. O.) lauroyl ethanolamide disodium salt, sulfosuccinic acid **polyoxyethylene** (2E. O.) monooleylamide disodium salt and the like; N-acylsulfonates such as sodium salt of cocoyl-N-methyltaurine, sodium salt of lauroyl-N-methyltaurine, triethanolamine. . . sodium palmityl sulfate, triethanolamine lauryl sulfate, triethanolamine myristyl sulfate triethanolamine palmityl sulfate and the like; ether sulfates such as triethanolamine **polyoxyethylene** (2-4E. O.) lauryl ether sulfate, sodium **polyoxyethylene** (2-4E. O.) myristyl ether sulfate, sodium **polyoxyethylene** (2-4E. O.) palmityl ether sulfate, triethanolamine **polyoxyethylene** (2-4E. O.) palmityl ether sulfate and the like; alkyl phosphates such as sodium lauryl phosphate, triethanolamine lauryl phosphate, sodium myristyl phosphate, triethanolamine myristyl phosphate, sodium oleyl phosphate, triethanolamine oleyl phosphate, sodium **polyoxyethylene** (2-4E. O.) lauryl ether phosphate, triethanolamine **polyoxyethylene** (2-4E. O.) lauryl ether phosphate, sodium **polyoxyethylene** (2-4E. O.) myristyl ether phosphate, triethanolamine **polyoxyethylene** (2-4E. O.) myristyl ether phosphate, sodium **polyoxyethylene** (2-4E. O.) oleyl ether phosphate, triethanolamine **polyoxyethylene** (2-4E. O.) oleyl ether phosphate, and the like.

SUMM [0026] As examples of the anti-inflammatory drug, there may be taken phenylbutazone, indomethacin, ibuprofen, ketoprofen, allantoin, guaiazulene, resorcin, hydrocortisone, prednisolone,

methylprednisolone, dexamethasone, triamcinolone, triamcinolone acetate, fludrocortisone, clobetasone, clobetasol and steroid esters thereof; ketal, acetal and hemiacetal derivative; flufenamic acid, bufexamac, naproxen, flurbiprofen, fenbufen, tenoxicam, piroxicam, mefenamic acid; salicylic acid, its derivative and salts thereof such as sodium salicylate, methyl salicylate, glycol salicylate and the like; D-panthenol, its derivative and.

SUMM . . . absorbers such as p-aminobenzoic acid, sodium p-aminobenzoate, ethyl p-aminobenzoate, butyl p-aminobenzoate, 2-ethylhexyl p-aminobenzoate, amyl p-dimethylaminobenzoate, glyceryl p-ibenzoate and the like; **salicylic acid** ultraviolet absorbers such as 2-ethylhexyl salicylate, triethanolamine salicylate, homomenthyl salicylate, dipropylene glycol salicylate, methyl salicylate, ethylene glycol salicylate, phenyl salicylate, . . .

SUMM . . . γ -linolenic acid, eicosapentaenoic acid, its derivative and salts thereof; organic acids selected from glycolic acid, succinic acid, lactic acid and **salicylic acid**, their derivatives and salts thereof; estradiol, its derivative and salts thereof; silk protein, its hydrolyzed product and derivatives thereof; hemoglobin. . .

SUMM [0031] As examples of the metal chelating agent, there may be taken malic acid, citric acid, **salicylic acid**, tartaric acid, gluconic acid, phytic acid, their derivatives and salts thereof; ethylenediamine-tetraacetic acid, its derivative and salts thereof; diethylenetriamine-pentaacetic acid, . . .

SUMM [0034] As examples of the solvent, there may be taken lower alcohols such as **ethanol** and the like, ethers, glycerins, liquid nonionic surfactants, liquid oily raw materials, other organic solvents, water, etc.

SUMM [0039] As examples of the percutaneous absorption promoters, there may be taken 2-pyrrolidone, 1-hexanol, 1-octanol, 1-decanol, 1-**menthol**, sodium lauryl sulfate, isopropyl myristate, n-hexyl acetate, oleic acid, etc.

SUMM [0040] As the steroid hormone, there may be taken 21-acetoxypregnenolone, alclometasone, algestone, almcinonide, beclomethasone, **betamethasone**, budesonide, chloroprednisone, clobetasone, clocortolone, clocprednol, **corticosterone**, cortisone, cortisol, deflazacort, desonide, diflorasone, diflucortolone, difluprednate, enoxolone, fluazacort, fluocloronide, flumethasone, flunisolide, fluocinolone acetate, fluocinolone, fluocortin butyl, fluocortolone, fluorometholone, fluperolone. . . acetate, fluprednisolone, flurandrenolide, formocortol, halcinonide, halometasone, halopredone acetate, hydrocortamate, hydrocortisone, hydrocortisone phosphate, hydrocortisone-21-succinate sodium salt, hydrocortisone tebutate, mazipredone, medrysone, meprednisone, **methylprednisolone**, mometasone furoate, paramethasone, prednicarbate, prednisolone-21-diethylaminoacetate, prednisolone sodium phosphate, prednisolone sodium succinate, prednisolone sodium-21-m-sulfobenzoate, prednisolone-21-stearoylglycolate, prednisolone tebutate, prednisolone-21-trimethylacetate, predonisone, prednival, prednylidene, prednylidene-21-diethylaminoacetate, tixocortol, **triamcinolone, triamcinolone acetate, triamcinolone benetonide, triamcinolone hexaacetonide, fluticasone**, etc.

DETD . . . skin induced by an organic acid. There were prepared Solution A of only glycolic acid dissolved in a 25% aqueous **ethanol** solution (the concentration of glycolic acid: 10%) and Solution B of glycolic acid and N, N'-diacetyl-L-cystine dimethyl ester dissolved in a 25% aqueous **ethanol** solution (the concentration of glycolic acid: 10%, the concentration of N, N'-diacetyl-L-cystine dimethyl ester: 10%). At that time, the pH. . .

DETD [0051] There were prepared Solution C of only glycolic acid dissolved in

a 25% aqueous **ethanol** solution (the concentration of glycolic acid: 20%) and Solution D of glycolic acid and N, N'-dioctanoyl-L-cystine dimethyl ester dissolved in a 25% aqueous **ethanol** solution (the concentration of glycolic acid: 20%, the concentration of N, N'-dioctanoyl-L-cystine dimethyl ester: 5%), and the function of N,.

DETD . . . of the skin induced an organic acid. There were prepared a solution of glycolic acid dissolved in a 25% aqueous **ethanol** solution (the concentration of glycolic acid: 10%) and a test solution of N, N'-diacetyl-L-cystine dimethyl ester dissolved in the same. . . .

DETD . . . and irritation induced by an organic acid. A test solution of N, N'-diacetyl-L-cystine dimethyl ester dissolved in a 25% aqueous **ethanol** solution (the concentration of N, N'-diacetyl-L-cystine dimethyl ester; 10%) was prepared. A solution of only the solvents was separately prepared. . . .

DETD [0066]

N,N'-di(n-lauroyl)-L-cystine dimethyl ester	2.0%
White Vaseline	25.0%
Stearyl alcohol	20.0%
Propylene glycol	12.0%
Polyoxyethylene hardened castor oil	4.0%
Glycerin monostearate	1.0%
Glycolic acid	1.0%
Trichloroacetic acid	1.0%
Antiseptic	q.s.
Perfume	q.s.
Purified water	Balance

DETD [0067]

N,N'-di(n-valeryl)-L-cystine dimethyl ester	3.0%
Glycolic acid	5.0%
Glycerin	3.0%
Sorbitol	2.0%
Polyoxyethylene (20) oleyl ether	1.0%
Ethanol	15.0%
Zinc p-phenol sulphonate	0.2%
Buffer	0.1%
Perfume	0.2%
Antiseptic	q.s.
Purified water	Balance

DETD [0068]

N,N'-di(n-propionyl)-L-cystine dimethyl ester	0.5%
Citric acid	1.0%
Urea	4.0%
Salicylic acid	2.0%
Lactic acid	2.0%
Glycerin	2.0%
Betaine	2.0%
Hyaluroinic acid	0.1%
Ethanol	15.0%
Buffer	0.1%
Perfume	0.2%
Antiseptic	q.s.
Purified water	Balance

DETD

N,N'-di(n-hexanoyl)-L-cystine dimethyl ester	0.5%
--	------

Lactic acid	0.1%
Fruit acid	0.1%
Glycerin	4.0%
Kaolin	1.0%
Caramine	0.7%
Camphor	0.2%
Ethanol	14.0%
Perfume	q.s.
Purified water	Balance
DETD [0070]	

N,N'-di (n-butyryl)-L-cystine dimethyl ester	1.0%	
Resorcinol	0.1%	
Kojic acid	1.0%	
Stearic acid	2.0%	
Polyoxyethylene (25) cetyl ether	3.0%	
Glyceryl monostearate	2.0%	
Octyl dodecanol	10.0%	
Cetanol	6.0%	
Reduced lanolin	4.0%	
Squalane	9.0%	
1,3-Butylene glycol	6.0%	
Polyethylene glycol (1500)	4.0%	
Antiseptic	q.s.	
Perfume	q.s.	
Purified water	Balance	
DETD . . . Solid paraffin		5.0%
Bees wax	10.0%	
Vaseline	15.0%	
Liquid paraffin	41.0%	
1,3-Butylene glycol	4.0%	
Glyceryl monostearate	2.0%	
Polyoxyethylene (20) sorbitan monolaurate	2.0%	
Borax	0.2%	
Antiseptic	q.s.	
Perfume	q.s.	
Antioxidant	q.s.	
Purified water	Balance	
DETD . . . 19.0%		
Stearic acid	5.0%	
Glyceryl monostearate	5.0%	
Sorbitan monostearate	12.0%	
Polyethylene sorbitan monostearate	38.0%	
Glycolic acid	1.0%	
Trichloroacetic acid	1.0%	
Chelating agent	q.s.	
Antiseptic	q.s.	
Perfume	q.s.	
Purified water	Balance	
DETD . . .		

N,N'-diacetyl-L-cystine	2.0%
N.sup.α-cocoylarginine ethyl ester DL-pyrrolidone	0.1%
carboxylate	
Retinol	0.1%
Bees wax	0.5%
Vaseline	2.0%
Glyceryl monostearate	1.0%
Polyethylene glycol monooleate	1.0%
Methyl polysiloxane	2.0%
Cetanol	1.0%

	Squalane	6.0%
	Carboxyvinyl polymer	0.5%
	1,3-Butylene glycol	4.0%
	Ethanol	5.0%
	Antiseptic	q.s.
	Perfume	q.s.
	Antioxidant	q.s.
	Purified water	Balance
DETD	. . . dimethyl ester 1.0%	
	Lactic acid	2.0%
	Stearyl alcohol	0.5%
	Hardened palm oil	3.0%
	Liquid paraffin	35.0%
	Dipropylene glycol	6.0%
	Polyethylene glycol (400)	4.0%
	Sorbitan sesquioleate	1.6%
	Polyoxyethylene (20) oleyl ether	2.4%
	Carboxyvinyl polymer	1.5%
	Potassium hydroxide	0.1%
	Chelating agent	q.s.
	Antiseptic	q.s.
	Perfume	q.s.
	Purified water	Balance
DETD	[0075]	

	N,N'-diacetyl-L-cystine diethyl ester	0.5%
	Liquid paraffin	12.0%
	Glyceryl tri(2-ethylhexanate)	50.0%
	Sorbit	10.0%
	Polyethylene glycol (400)	5.0%
	Lactic acid	1.0%
	Acylmethyl taurine	4.0%
	Polyoxyethylene (20) isocetyl ether	10.0%
	Perfume	q.s.
	Antiseptic	q.s.
	Purified water	Balance
DETD	[0076]	

	N,N'-di(n-heptanoyl)-L-cystine dimethyl ester	0.5%
	Fruit acid	0.5%
	Dipropylene glycol	5.0%
	Polyethylene glycol (400)	5.0%
	Ethanol	10.0%
	Carboxyvinyl polymer	0.5%
	Sodium alginate	0.5%
	Potassium hydroxide	0.2%
	Polyoxyethylene (20) sorbitan monostearate	1.0%
	Sorbitan monooleate	0.5%
	Oleyl alcohol	0.5%
	Placenta extract	0.2%
	dl-tocopherol acetate	0.2%
	Perfume	q.s.
	Antiseptic	q.s.
DETD	[0077]	

	N,N'-diacetyl-L-cystine diisopropyl ester	3.0%
	Isopropanol	2.0%
	Polyvinyl alcohol	15.0%
	Carboxymethylcellulose	5.0%

1,3-Butylene glycol	5.0%
Ethanol	12.0%
Sodium alginate	0.5%
Polyoxyethylene (20) oleyl ether	0.5%
Perfume	q.s.
Antiseptic	q.s.
Buffer	q.s.
Purified water	Balance
DETD [0078]	

N,N'-di(n-octanoyl)-L-cystine dimethyl ester	5.0%
Salicylic acid	0.5%
Liquid paraffin	10.0%
Polyoxyethylene (20) sorbitan monooleate	3.5%
Propylene glycol	3.0%
Titanium oxide	9.0%
Kaolin	24.0%
Talc	42.0%
Coloring pigment	3.0%
Perfume. . .	
DETD [0080]	

N,N'-di (n-butyryl)-L-cystine diamide	0.5%
Triethylamine N-lauroylglutamate	25.0%
Triethanolamine laurate	5.0%
Polyoxyethylene (4) polyoxypropylene (11) butyl ether	5.0%
Ethanol	3.0%
Perfume	q.s.
Antiseptic	q.s.
Purified water	Balance
DETD [0081]	

N,N'-dioctanoyl-L-cystine dimethyl ester	0.1%
Triethanolamine polyoxyethylene (3) lauryl ether sulfate	3.0%
Sodium polyoxyethylene (3) lauryl ether sulfate	6.0%
Sodium lauryl sulfate	1.5%
Lauric acid diethanolamide	3.0%
DETD [0082]	

N,N'-diacetyl-L-cystine dimethyl ester	1.0%
Lactic acid	0.02%
Oleyl alcohol	0.2%
Liquid paraffin	0.5%
Ethanol	5.0%
Sorbitol	4.0%
Polyoxyethylene (20) lauryl ether	2.5%
Sorbitan monolaurate	0.5%
Pigment	0.1%
Antiseptic	0.1%
Perfume	0.1%
Purified water	Balance
DETD [0083]	

N,N'-diacetyl-L-cystine dimethyl ester	2.0%
Ethanol	70.0%

Benzalkonium chloride	1.0%
Chlorhexidine gluconate	0.5%
2-Lauryl-N-carboxymethyl-N-hydroxyethylimidazolium betaine	2.0%
Sodium DL-pyrrolidone carboxylate	5.0%
N.sup.alpha.-cocoylarginine ethyl ester DL-pyrrolidone	1.0%

DETD [0084]

N,N'-dioctanoyl-L-cystine dimethyl ester	0.1%
Lauryl dimethylaminoacetic acid betaine	5.0%
Sodium chloride	1.0%
Benzalkonium chloride	0.1%
Stearyl trimethyl ammonium chloride	2.0%
Boric acid	0.5%
Borax	0.1%
EDTA	0.1%
Purified water	Balance

IT 50-21-5, Lactic acid, biological studies 67-63-0, Isopropanol, biological studies 69-72-7, Salicylic acid, biological studies 77-92-9, Citric acid, biological studies 79-14-1, Glycolic acid, biological studies 112-03-8, Stearyltrimethylammonium chloride 151-21-3, Sodium laurylsulfate, biological studies 683-10-3, Lauryldimethylamino acetate betaine 2224-49-9, Triethanolamine laurate 5545-17-5, N,N'-Diacetyl-L-cystine 9004-82-4, Polyoxyethylene lauryl ether sulfate sodium salt 18472-51-0, Chlorhexidine gluconate 25859-09-0, Diethanolamine laurate 27028-82-6, Polyoxyethylene lauryl ether sulfate triethanolamine salt 32381-28-5 41489-26-3D, N.alpha.-cocoyl derivs. 53576-49-1 54571-67-4 121972-22-3 139416-22-1 139416-28-7 139416-29-8 193697-38-0 263553-64-6 263553-65-7 263553-66-8 460040-30-6
(skin compns. containing cystine derivs., and chemical peeling agents, bactericides, and/or surfactants)

L130 ANSWER 2 OF 8 USPATFULL on STN

AN 2003:180375 USPATFULL

TI Formulations and methods for reducing skin irritation

IN Hahn, Gary S., Cardiff by the Sea, CA, UNITED STATES

Thueson, David O., Poway, CA, UNITED STATES

PI US 2003124202 A1 20030703

AI US 2002-189344 A1 20020703 (10)

RLI Continuation of Ser. No. US 2001-33194, filed on 24 Oct 2001, PENDING
Continuation of Ser. No. US 2001-853282, filed on 11 May 2001, PENDING
Continuation of Ser. No. US 2000-685992, filed on 10 Oct 2000, ABANDONED
Continuation of Ser. No. US 1997-860993, filed on 23 Jun 1997, GRANTED,
Pat. No. US 6139850 Continuation-in-part of Ser. No. US 1994-362100,
filed on 21 Dec 1994, GRANTED, Pat. No. US 5716625 Continuation-in-part
of Ser. No. WO 1995-US16985, filed on 21 Dec 1995, PENDING

DT Utility

FS APPLICATION

LREP Cosmederm Technologies, LLC, 4370 La Jolla Village Drive, Suite 960, San Diego, CA, 92122

CLMN Number of Claims: 92

ECL Exemplary Claim: 1

DRWN 13 Drawing Page(s)

LN.CNT 1619

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Compositions and methods are provided for inhibiting skin irritations attributable to chemical irritants or environmental conditions by the application of anti-irritant amounts of aqueous-soluble divalent strontium cation.

AN 2003:180375 USPATFULL

SUMM . . . which the active drug ingredients are formulated may also produce irritation in sensitive people, especially in drugs such as topical **corticosteroids**.

SUMM . . . as retinoids (e.g. tretinoin, retinol and retinal), carboxylic acids including α -hydroxy acids (e.g. lactic acid, glycolic acid), β -hydroxy acids (e.g. **salicylic acid**), α -keto acids, acetic acid and **trichloroacetic acid**, 1-pyrrolidone-5-carboxylic acid, capryloyl **salicylic acid**, α -hydroxy decanoic acid, α -hydroxy octanoic acid, gluconolactone, methoxypropyl gluconamide, oxalic acid, malic acid, tartaric acid, mandelic acid, benzylic acid, gluconic. . .

SUMM . . . the irritation caused by such products. Common exfolients include α - and β -hydroxy carboxylic acids such as lactic acid, glycolic acid, **salicylic acid** and the like, α -keto acids such as pyruvic acid, as well as assorted compounds such as acetic acid and **trichloroacetic acid**, 1-pyrrolidone-5-carboxylic acid, capryloyl **salicylic acid**, α -hydroxy decanoic acid, α -hydroxy octanoic acid, gluconolactone, methoxypropyl gluconamide, oxalic acid, malic acid, tartaric acid, mandelic acid, benzylic acid, gluconic. . .

SUMM . . . topical application of the hydroxy acid skin irritant lactic acid as well as the skin irritants glycolic acid, capsaicin, capryloyl **salicylic acid**, benzoyl peroxide, and post-shaving-applied seawater. Formulations containing the strontium cation are useful in suppressing a wide range of topical-product-induced irritation. . .

SUMM . . . example in which the vehicle has a density of 0.93 g/ml (as in a 50:50 [by volume] mixture of 95% **ethyl alcohol** and water) and the cation component is incorporated in the form of strontium nitrate (formula weight 212), represent molarity concentration. . .

SUMM . . . components) as water; organic solvents such as alcohols (particularly lower alcohols readily capable of evaporating from the skin such as **ethanol**), glycols (such as glycerin), aliphatic alcohols (such as lanolin); mixtures of water and organic solvents (such as water and alcohol),. . .

DETD . . . skin irritation caused by certain severe skin irritants, including particularly lactic acid and glycolic acid (which are hydroxy acids), capryloyl **salicylic acid** (a β -hydroxy acid ester) and capsaicin (an isolate from cayenne and paprika known for its skin-irritating properties). The trials were. . .

DETD . . . of the subjects, In the majority of the tests, the irritant composition was 7.5% lactic acid dissolved in a 10% **ethanol** -in-water solution.

DETD . . . of the irritant composition. Controls were performed by applying corresponding formulation(s) (pretreatment and/or skin-irritant composition) with an equimolar amount of **sodium chloride** to a contralateral portion of the subject's skin. Typically, the test materials were applied to the face of the subject.

DETD . . . Chloride SrCl.sub.2 VIS DIFFERENCE 20

Strontium Nitrate Sr(NO.sub.3).sub.2 VIS DIFFERENCE 56

Strontium Acetate Sr(CH.sub.3CO.sub.2).sub.2 VIS DIFFERENCE 46

TIME ZERO TESTS

Strontium Chloride SrCl.sub.2 10% EtOH 58

Strontium Nitrate Sr(NO.sub.3).sub.2 10% EtOH 64

DETD . . . test compounds of the invention were formulated in Elizabeth Arden "Visible Difference Refining Toner", with the Toner mixed with equimolar **sodium chloride** serving as the control. The test solutions (and control) were provided in coded vials for application to either the right. . .

DETD [0079] Following a protocol parallel to that of the lactic acid irritant trials described above, glycolic acid (6.0% in 10% **ethanol** -in-water) was applied as a skin irritant to subject panels. Strontium

nitrate was co-administered as an anti-irritant (time zero testing), and. . .

DETD . . . subject females. The control solution was Vaseline Smooth Legs and Feet Lotion (containing water, lactic acid (5%), glycerin, isopropyl palmitate, PEG-40 stearate, cetyl alcohol, potassium hydroxide, steareth-2, magnesium aluminum silicate, lecithin, soya sterol, tocopheryl acetate, tetinyl palmitate, dimethicone, menthol, camphor, stearic acid, laureth-7, xanthan gum, polyacrylamide, C13-14 isoparaffin, corn oil, fragrance, DMDM hydantoin, iodopropynyl butylcarmamate, disodium EDTA, PG, and. . .

DETD . . . cyclomethicone (Dow Corning, "DC344"), 7.5 ml cyclomethicone/dimethiconol (Dow Corning, "DC1401"), 7.5 ml cyclomethicone/dimethicone copolyol (Dow Corning, "DC3225C") and 8 ml PEG-8 and blended for 2-3 minutes. Imidizolidinyl urea (0.5%) was added as a preservative. A clear, thick gel resulted (50 ml)..

DETD . . . of the salt with Mary Kay Revival Serum (with 15% lactic acid) and L'Oreal Vichy Novaetia Cream (with 2% capryloyl salicylic acid), respectively.

CLM What is claimed is:

15. The composition of claim 12 wherein said irritant ingredient comprises salicylic acid or a salt thereof.

16. The composition of claim 12 wherein said irritant ingredient comprises a combination of lactic acid and salicylic acid, or salts thereof.

17. The composition of claim 12 wherein said irritant ingredient comprises capryloyl salicylic acid or a salt thereof.

23. The composition of claim 12 wherein said irritant ingredient comprises trichloroacetic acid of a salt thereof.

IT 50-21-5, Lactic acid, biological studies 57-13-6, Urea, biological studies 58-08-2, Caffeine, biological studies 64-19-7, Acetic acid, biological studies 68-26-8, Retinol 69-72-7, Salicylic acid, biological studies 76-03-9, Trichloroacetic acid, biological studies 76-93-7, biological studies 77-92-9, Citric acid, biological studies 79-14-1, Glycolic acid, biological studies 87-69-4, Tartaric acid, biological studies 90-64-2, Mandelic acid 90-80-2, Gluconolactone 94-36-0, Benzoyl peroxide, biological studies 97-59-6, Allantoin 98-79-3 108-95-2, Phenol, biological studies 116-31-4, Retinal 127-17-3, Pyruvic acid, biological studies 144-62-7, Oxalic acid, biological studies 302-79-4, Tretinoin 515-69-5, α -Bisabolol 526-95-4, Gluconic acid 543-94-2, Strontium acetate 617-73-2, α -Hydroxyoctanoic acid 1405-86-3, Glycyrrhizic acid 5393-81-7, α -Hydroxydecanoic acid 6915-15-7, Malic acid 7440-24-6, Strontium, biological studies 7759-02-6, Strontium sulfate 10042-76-9, Strontium nitrate 10476-85-4, Strontium chloride 70424-62-3 126094-21-1
(strontium cation formulations for reducing skin irritation)

L130 ANSWER 3 OF 8 USPATFULL on STN

AN 2002:246386 USPATFULL

TI Method of applying an adhesive composition over a bioactive polymerization initiator or accelerator

IN Narang, Upvan, Raleigh, NC, United States
Hedgpeth, Daniel L., Raleigh, NC, United States
Szabo, Gabriel N., Raleigh, NC, United States
Badejo, Ibraheem T., Morrisville, NC, United States
Barefoot, Joe B., Raleigh, NC, United States

PA Closure Medical Corporation, Raleigh, NC, United States (U.S. corporation)

PI US 6455064 B1 20020924
 AI US 1999-430176 19991029 (9) <--
 RLI Continuation-in-part of Ser. No. US 1998-69875, filed on 30 Apr 1998
 DT Utility
 FS GRANTED
 EXNAM Primary Examiner: Dees, Jose' G.; Assistant Examiner: Williamson, Michael A.
 LREP Oliff & Berridge, PLC
 CLMN Number of Claims: 47
 ECL Exemplary Claim: 1
 DRWN 0 Drawing Figure(s); 0 Drawing Page(s)
 LN.CNT 1181

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A composition comprising a polymerizable adhesive monomer is applied over a biologically active initiator or accelerator for polymerization of the monomer. The biologically active initiator or accelerator is a medicament that provides a desired medical or therapeutic activity as well as enhancing polymerization of the adhesive.

AN 2002:246386 USPATFULL

AI US 1999-430176 19991029 (9) <--

SUMM . . . bioactive agents is also disclosed in: Miles et al., Oral Surgery, Oral Medicine, Oral Pathology, Vol. 75, No. 3, 397-402 (using **triamcinolone** acetonide (Kenalog) or chlorhexidine digluconate (Peridex) as the bioactive agent); and Kaufman, R. S., The Laryngoscope, 1974, 793-804 (using **dexamethasone** sodium phosphate (Decadron) as the bioactive agent).

SUMM . . . not to affect the cure rate or the bond strength of the glue layer. They include thymol, chlorothymol, benzoic acid, p-**hydroxybenzoate** alkyl esters, 4- and 6-phenyl-2-chlorophenyl, carvocrol, hexachlorophene, nitroforans, allicin, 2-phenylphenol, boric acid, mercurials, and such antibiotics as Bacitracin and Griseofulvin, quaternary ammonium halides such as n-alkyldirnethylbenzylammonium chloride, cetyl pyridinium bromide, 5-methyl-2-isopropyl-cyclohexanol, 2-bornanone, cineole, safrole, bornyl chloride, 2-phenoxyethanol, benzylalcohol and **ethanol**. The biocides are applied to human fingernails, then covered by solutions comprising cyanoacrylate adhesive. The biocides are applied to the. . .

SUMM . . . mixtures thereof. Preferred plasticizers are tributyl citrate and acetyl tributyl citrate. In embodiments, suitable plasticizers include polymeric plasticizers, such as **polyethylene glycol** (PEG) esters and capped PEG esters or ethers, polyester glutarates and polyester adipates.

SUMM . . . 2,4-dinitrophenol (pK.sub.a4.0), formic acid (pK.sub.a3.7), nitrous acid (pK.sub.a3.3), hydrofluoric acid (pK.sub.a3.2), chloroacetic acid (pK.sub.a2.9), phosphoric acid (pK.sub.a2.2), dichloroacetic acid (pK.sub.a1.3), **trichloroacetic acid** (pK.sub.a0.7), 2,4,6-trinitrophenol (picric acid) (pK.sub.a0.3), trifluoroacetic acid (pK.sub.a0.2), sulfuric acid (pK.sub.a3.0), sulfurous acid, and mixtures thereof. In embodiments, the amount. . .

SUMM . . . (Florida Chemical Co.), cold pressed lime oil (Florida Chemical Co.), cucumber distillate (Florida Chemical Co.), honey distillate (Florida Chemical Co.), **menthol** (Aldrich), alkyl salicylates such as methyl salicylate (Lorann Oils or Aldrich), monosodium glutamate, spearmint, wintergreen, cinnamon, citrus, cherry, apple, peppermint, . . .

SUMM . . . are not limited to, waxes, such as carnauba, petroleum and carbowax; gels, such as gelatin, hydroxypropyl methylcellulose, carboxymethylcellulose, and hydroxy-gels; **polyethylene glycol**; polysorbate; agar; povidone; sodium stearate; starch; powdered sugar; high fructose corn syrup; fructose; glycerin; hydrogenated glucose syrup; sorbitol; mannitol; sucrose; cellulose acetate phthalate; dextrose; **polyvinyl alcohol**;

mixtures thereof; and the like.

SUMM Suitable alcohols include phenols, 1,4-butanediol, d-sorbitol, and **polyvinyl alcohol**.

CLM What is claimed is:

. . . pressed valencia orange oil, cold pressed grapefruit oil, cold pressed lemon oil, cold pressed lime oil, cucumber distillate, honey distillate, **menthol**, alkyl salicylates, monosodium glutamate, spearmint, wintergreen, cinnamon, citrus, cherry, apple, peppermint, peppermint oil, peppermint spirit, vanillin, thymol, and ethyl vanillin.

IT 69-72-7D, Salicylic acid, alkyl esters 89-78-1, Menthol 89-83-8, Thymol 100-52-7, Benzaldehyde, biological studies 104-46-1, Anethole 121-32-4, Ethyl vanillin 121-33-5, Vanillin 142-47-2, Monosodium glutamate 25916-47-6, Poly(acrylic acid), zinc salt 336804-71-8, Poly(cyanoacrylic acid), zinc salt (medical adhesive compns. containing monomers and bioactive polymerization initiators or accelerators)

L130 ANSWER 4 OF 8 USPATFULL on STN

AN 2000:174129 USPATFULL

TI Preparation for the application of agents in mini-droplets

IN Cevc, Gregor, Heimstetten, Germany, Federal Republic of

PA Idea AG, Munich, Germany, Federal Republic of (non-U.S. corporation)

PI US 6165500 20001226 <--

AI US 1992-844664 19920408 (7) <--

PRAI DE 1990-4026834 19900824 <--

DE 1990-4026833 19900824 <--

DE 1991-4107153 19910306 <--

WO 1991-EP1596 19910822 <--

DT Utility

FS Granted

EXNAM Primary Examiner: Kishore, Gollamudi S.

LREP Davidson, Davidson & Kappel, LLC

CLMN Number of Claims: 35

ECL Exemplary Claim: 1

DRWN 31 Drawing Figure(s); 21 Drawing Page(s)

LN.CNT 4336

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention relates to a preparation for the application of agents in the form of minuscule droplets of fluid, in particular provided with membrane-like structures consisting of one or several layers of amphiphilic molecules, or an amphiphilic carrier substance, in particular for transporting the agent into and through natural barriers such as skin and similar materials. The preparation contains a concentration of edge active substances which amounts to up to 99 mol-% of the agent concentration which is required for the induction of droplet solubilization. Such preparations are suitable, for example, for the non-invasive applications of antidiabetics, in particular of insulin. The invention, moreover, relates to the methods for the preparation of such formulations.

AN 2000:174129 USPATFULL

PI US 6165500 20001226 <--

AI US 1992-844664 19920408 (7) <--

PRAI DE 1990-4026834 19900824 <--

PRAI DE 1990-4026833 19900824 <--

PRAI DE 1991-4107153 19910306 <--

PRAI WO 1991-EP1596 19910822 <--

SUMM . . . prolonged drug action but has not increased the skin-penetration capability of the drug itself. Through massive use of penetration enhancers (**polyethylene glycol** and fatty acids) and of lipid vesicles, Gesztes und Mezei (1988, Anesth. Analg. 67, 1079-1081) have succeeded in inducing local. . .

DETD . . . are to some extent edge active only in certain concentration

ranges encompass simple, especially short chain, alcohols, such as methanol, **ethanol**, n-propanol, 2-propen-1-ol(allyl alcohol), n-butanol, 2-buten-1-ol, n-pentanol (amyl alcohol), n-hexanol, n-heptanol, n-octanol and n-decanol; furthermore, iso-propanol, iso-butanol or iso-pentanol. Higher alcohols are even. . . the present purpose as well as cyclic alcohols, such as benzyl alcohol, cyclopentanol, cyclohexanol, 3-, 4-, 5-cyclohexanol, cyclohexyl alcohol, aryl-alcohols, such as phenyl-**ethanol**, etc.

DETD Sorbitol is one possible example of residue Z. (X.sub.i --Y.sub.j) can be a polyene, polyoxyalkene, such as **polyoxyethylene**, polyalcohol, such as polyglycol, or polyether. (X.sub.i --Y.sub.j) mainly contain 1-20 and very frequently 2-10 units, e.g. in ethylene glycol, di- and triglycol (oligoglycol) or **polyethylene glycol**.

DETD . . . surfactants of the ether-type which are suitable for the present purpose are the substances of the Myrj trademark, such as **polyoxyethylene**(8)-stearate (Myrj45), **polyoxyethylene**(20)-stearate (Myrj49), **polyoxyethylene**(30)-stearate (Myrj51), **polyoxyethylene**(40)-stearate (Myrj52), **polyoxyethylene**(50)-stearate (Myrj53), **polyoxyethylene**(100)-stearate (Myrj59), etc. Further products of these classes are sold under the trademark Cirrasol ALN; common **polyoxyethylene**-alkylamides are e.g. surfactants of the trademark Atplus.

DETD . . . R.sub.3 and R.sub.4 are frequently of the alkoxy- or alkenoxy-, and even more commonly of the polyene-, polyoxyalkene-, such as **polyoxyethylene**-, polyalcohol-, such as polyglycol-, or polyether type. Some of these chains can be apolar, corresponding to e.g. an acyl-, alkyl-, . . .

DETD Chains in the substances of TWEEN type are very frequently of the **polyoxyethylene** class. They mainly contain one terminal hydrogen atom and more rarely a methoxy group. One of the **polyoxyethylene** chains, however, contains a hydrophobic residue which preferably corresponds to an acyl-, alkyl-, alkenyl-, hydroxyalkyl-, hydroxyalkenyl- or hydroxyacyl-chain with 4-24, . . .

DETD . . . this case is cationic, in order to ensure that the whole molecule is zwitterionic. Most frequently, ammonio-alkyl derivatives, such as **ethanol**-, propanol-, butanol-, pentanolamine, hexanolamine, heptanolamine or octanolamine, N-methyl-, N,N-dimethyl, or N,N,N-trimethyl-ammonio-alkyl, N-ethyl-, N,N-diethyl, or N,N,N-triethyl-amino-alkyl, unequal N-alkyles, such as N,N-methyl-ethyl-ammonio-alkyl, . . .

DETD . . . PX, Thesit), nonyl-glucoside, octaethylene-glycol-isotridecylether (Genapol X-080), octaethylene-dodecyl-ether, octanoyl-N-methyl-glucamide (MEGA-8), 3-(octyldimethylammonio)-propanesulfonate (Zwittergent 3-08), octyl-glucoside, octylthiogluconate, entadecaethylene-isotridecyl-ether (Genapol X-150), polyethylene-polypropylene-glycol (Pluronic F-127), **polyoxyethylene**-sorbitane-monolaurate (Tween 20), **polyoxyethylene**-sorbitane-monoleate (Tween 80), taurodeoxycholate-sodium salt, taurocholate-sodium salt, 3-(tetradecyldimethylammonio)-propane-sulfonate (Zwittergent 3-14), etc.

DETD . . . salts (sodium dodecylsulfate, Duponol C, SDS, Texapon K12), N-hexadecyl-sulfobetaine (Zwittergent 3-16), nonaethylene-glycol-octyl-phenyl-ether (NP-40, Nonidet P-40), nonaethylene-dodecyl-ether, octaethylene-glycol-isotridecyl-ether (Genapol X-080), octaethylene-dodecyl-ether, **polyethylene glycol** -20-sorbitane-monolaurate (Tween 20), **polyethylene glycol**-20-sorbitane-monostearate (Tween 60), **polyethylene glycol**-20-sorbitane-monoleate (Tween 80), polyhydroxyethylenecetylstearyl ether (Cetomacrogol, Cremophor 0, Eumulgin, C 1000) polyhydroxyethylene-4-lauryl ether (Brij 30), polyhydroxyethylene-23-lauryl ether (Brij 35), polyhydroxyethylene-8-stearate (Myrj 45, Cremophor AP), . . . oil (Cremophor RH 40,

Cremophor RH 60) polyethoxylated plant oils (Lebrafils), sorbitane-monolaurate (Arlacel 20, Span 20), taurodeoxycholate salts, taurocholate salts, **polyethylene glycol** -20-sorbitane-palmitate (Tween 40), Myrj 49 and **polyethylene glycol** derivatives of ricinols, etc.

- DETD . . . phenylbutazone-derivatives (such as 3,5 pyrazolidine dion), pherazone, piroxicam, propoxyphene, propyphenazon, pyrazol- and phenazone-derivatives (aminophenazone, metamizole, monophenylbutazone, oxyphenebutazone, phenylbutazone or phenazonesalyzilate), **salicylic acid**-derivatives, sulfasalazine, tilidine; acetylsalicylic acid, ethylmorphine, alclofenac, alphaprodine, aminophenazone, anileridine, azapropazone, benfotiamine, benorilate, benzydamine, cetobemidone, chlorophenesincarbamate, chlorothenoazine, codeine, dextromoramide, dextro-propoxyphene, ethoheptazine, . . .
- DETD . . . as aminophenazole, bemegride, caffeine, doxapram, ephedrine, prolintane, or nialamide and tranylcypromine; but also vitamins, plant extracts from semen colae, camphor, **menthol**;
- DETD . . . least one substance from the class of antiallergics: e.g. agents from the globuline family, corticoids or antihistaminics (such as beclometasone-, **betametasonecortisone**-, **dexametasone** -derivatives, etc.) as well as bamipinacetate, buclizine, clemastine, clemizole, cromoglicinic acid, cyproheptadine, diflucorolonvalerate, dimetotiazine, diphenhydramine, diphenylpyraline, ephedrine, fluocinolane, histapyrrodine, isothipendyle, methadilazine, . . .
- DETD . . . the antiasthmatics and/or bronchospasmolytics, such as amiodarone, carbutole, fenoterol, orciprenalin, sotalol, or theophylline-derivatives, as well as corticoids (such as beclomethasone, **dexamethasone**, hydrocortisone, prednisolone), frequently in combination with purines;
- DETD . . . or vitamins, etc., are preferred for this purpose, as well as antiphlogistics, such as quinine, nicotinic acid-, nonylic acid-, or **salicylic acid**-derivatives, meproamate, etc.;
- DETD at least one glucocorticoid, such as beclomethason, **betamethason**, clocortolone, cloprednol, cortison, **dexamethason** (e.g. as a **dexamethasonephosphate**), fludrocortison, fludroxcortide, flumetason, fluocinoloneacetone, fluocinonide, fluocortolon (e.g. as a fluocortoloncapronate or fluocortolontrimethylacetate), fluorometholon, fluprednidenacetate, hydrocortison (also as a hydrocortison-21-acetate, hydrocortison-21-phosphate, etc.), paramethason, prednisolon (e.g. in the form of **methylprednisolon**, prednisolon-21-phosphate, prednisolon-21-sulfobenzoate, etc.), prednison, prednylidene, pregnenolon, **triamcinolon**, **triamcinolonacetone**, etc.;
- DETD . . . (Methyl 4-methylpyrrole 2-carboxylate) cis-13-octadecenal 13-octadecyn-1-ol, 2-(phenyl)ethyl propionate(phenylethanol propanoate), propyl cyclohexylacetate, cis-9,trans-11-tetradecadienol ([Z,E]-9,11-TDDOL), cis-9, trans-11-tetradecadienyl acetate ([Z,E]-9,11-TDDA), cis-9, trans-12-tetradecadienyl acetate ([Z,E]-9,12-TDDA), **trichloroacetic acid** esters, cis-9-tricosene, undecanal, etc.;
- DETD . . . myeloperoxidase (1.11.1.7), peroxidase (1.11.1.7), glutathione peroxidase (1.11.1.9), chloroperoxidase (1.11.1.10), lipoxidase (1.13.1.12), protococatechuate 3,4-dioxygenase (1.13.11.3), luciferase (glow-worm) (1.13.12.7), salicylate hydroxylase (1.14.13.7), p-**hydroxybenzoate** hydroxylase (1.14.13.2), luciferase (bacterial) (1.14.14.3), phenylalanine hydroxylase (1.14.16.1), dopamine-beta-hydroxylase (1.14.17.1), tyrosinase (1.14.18.1), superoxide dismutase (1.15.1.1), ferredoxine-NADP reductase (1.18.1.2), etc.. Transferases, . . .
- DETD . . . halogenated, aliphatic, cycloaliphatic, aromatic or aromatic-aliphatic hydrocarbons, such as benzol, toluol, methylene chloride or chloroform, alcohols, such as methanol or **ethanol**, propanediol, erithritol, short-chain alkane carboxylic acid esters, such

as acetic acid alkylesters, such as diethylether, dioxan or tetrahydrofuran, or. . .

DETD

250-372 mg phosphatidylcholine from soy-bean (+95% = PC)

187-34.9 mg

oleic acid (+99%)

0.312-0.465 ml

ethanol, absolute

10 mM

Hepes

DETD

322.6-372 mg

phosphatidylcholine from soy-bean (+95% = PC)

96.8-34.9 mg

oleic acid (+99%)

0.403-0.465 ml

ethanol, absolute

10 mM

Hepes

130 mM

NaCl, p.a.

DETD

. . . phosphatidylcholine from soy-bean (+95% = PC)

20.5-22.2 mg

phosphatidylglycerol from egg PC (puriss.,

Na-salt, = PG)

44.9-26.1 µl

oleic acid (+99%)

0.165-0.178 ml

ethanol, absolute

4.5 ml

Hepes, 10 mM

DETD

301.3-335.4 mg

phosphatidylcholine from soy-bean (+95% = PC)

123.3-80.8 µl

Tween 80 (puriss.)

0.38-0.42 ml

ethanol, absolute

4.5 ml

phosphate buffer, isotonic, sterile

DETD

193-361 mg

phosphatidylcholine from soy-bean (grade I, S100)

207.2-38.8 mg

Na-cholate, puriss.

4.5 ml

phosphate buffer (isotonic with a physiologic solution)

ethanol, absolute

DETD

0.5 ml of a hot solution of S100 in **ethanol** (2/1, M/V) are mixed with sufficient amounts of bile acid salts which give rise to a concentration series with increasing. . .

DETD

121.2-418.3 mg

phosphatidylcholine from soy-bean (Grade I,

PC) 378.8-81.7 mg

Triton X-100

4.5 ml

0.9% **NaCl** solution in water

DETD

A 10% PC-suspension in isotonic solution of **sodium chloride** is homogenized at 22° C. until the mean size of lipid vesicles is approx. 400 nm. This suspension is then. . .

DETD

101.6-227 mg phosphatidylcholine from soy-bean

148.4-22.2 mg octyl-glucopyranoside (β -octylglucoside),
 puriss. 9.85 ml
 phosphate buffer, pH 7.3
 ethanol, absolute

DETD Phosphatidylcholine in **ethanol** (50%) and octylglucopyranoside
 were mixed in different relative ratios in order to prepare a
 concentration series with increasing L/S values. . . .

DETD

84.2 to 25 mg

 phosphatidylcholine from soy-bean 80%

75 kBq Gibberellin A4, 3H-labelled

15.8 to 75 mg

polyoxyethylene (23)-laurylether (Brij 35)

1 ml water

ethanol, absolute

DETD . . . dipalmitoylphosphatidylcholine in a chloroform solution. The
 resulting lipid mixture is dried and then dissolved in 30 microliters of
 warm, absolute **ethanol**. This solution is then mixed with 0.32
 ml of a buffer solution (phosphate buffer, 10 mM, 0.9% NaCl);
 this corresponds to a lipid/surfactant ratio of 4/1. The resulting
 suspension is thoroughly mixed and subsequently filtered through filters
 with. . . .

DETD

88 mg phosphatidylcholine from soy-bean (purity higher
 than 95%, PC)

75 kBq insulin, tritium labelled

12 mg deoxycholate, Na-salt, p.a.

100 ml **ethanol**, absolute

0.9 ml isotonic salt solution

DETD 100 mg of PC dissolved in 100 ml of warm **ethanol**, or a
 corresponding PC/deoxycholate solution (L/S=4.5), are mixed with 0.9 ml
 of an isotonic salt solution (suspensions A and B,

DETD

386 mg phosphatidylcholine from soy-bean
 (purity > 95%)

58.5 mg sodium-cholate (L/S = 3.5)

500 ml **ethanol** (96%)

2.25 ml 0.9% NaCl solution (per inject.)

2.25 ml Actrapid HM 40 (corresponds to 90 I.U. of
 recombinant human insulin)

DETD . . . mixture of aqueous salt solution and human recombinant insulin
 (with 6.75 mg m-cresole) is mixed with a lipid solution in
 ethanol. The resulting, opaque suspension is aged over night. 12
 hours later, this suspension is pressed through a sterile filter
 (Anodisc,

DETD

956 mg phosphatidylcholine from soy-bean (+95%)

0-26 mg sodium-deoxycholate

1 mg prostaglandine E1

1 ml **ethanol** absolute

50 ml 0.9% NaCl solution (per inject.)

DETD 1 ml of **ethanol** is pipetted into a glass flask with 1 mg of
 prostaglandine. After thorough mixing, the resulting prostaglandine
 solution is transferred. . . . with 6 ml of an isotonic salt solution.
 The prostaglandine containing flask is washed twice with 2 ml of 0.9%
 NaCl and mixed with the original lipid suspension. The sample is
 then divided into 5 parts; into individual aliquots sodium-desoxycholate
 is. . . .

DETD

79.4 mg; 88.5 mg
 phosphatidylcholine from soy-bean (+95%)
 20.6 mg, 11.5 mg
 sodium-deoxycholate
 10 µg hydrocortison
 0.1 ml **ethanol** absolute
 1 ml phosphate buffer, physiological

DETD

256.4-447 mg
 phosphatidylcholine from soy-bean (+95% PC)
 243.6-53.1 mg
 Brij 96
 0.26-0.45 ml
 ethanol, absolute
 4.5 ml phosphate buffer, pH 6.5. 10 mM

DETD

202.0-413 mg
 phosphatidylcholine from soy-bean (+95% = PC)
 298.0-87.0 mg
 Myrj 49
 0.26-0.45 ml
 ethanol, absolute
 4.5 ml phosphate buffer, pH 6.5. 10 mM

DETD

144.9 mg phosphatidylcholine from soy-bean
 24.8 mg desoxycholate, Na-salt
 1.45 ml Actrapid HM 100 (145 I.U.)
 0.16 ml **ethanol**, absolute

DETD Appropriate quantities of both lipids are dissolved in corresponding amounts of **ethanol** and mixed with a standard solution of insulin. 12 hours later, the crude carrier suspension is homogenized by means of. . . .

DETD . . . alkylsulfate-salts, cholate-, deoxycholate-, glycodeoxycholate-, taurodeoxycholate-salts, dodecyl-dimethyl-aminoxide, decanoyl- or dodecanoyl-N-methylglucamide (MEGA 10, MEGA 12), N-dodecyl-N,N-dimethylglycine, 3-(hexadecyldimethylammonio)-propanesulfonate, N-hexadecylsulfobetaine, nonaethyleneglycol-octylphenylether, nonaethylene-dodecylether, octaethyleneglycol-isotridecylether, octaethylene-dodecylether, **polyethylene glycol**-20-sorbitane-monolaurate (Tween 20), **polyethylene glycol**-20-sorbitane-monooleate (Tween 80), polyhydroxyethylenecetylstearylether (Cetomacrogol, Cremophor O, Eumulgin, C 1000) polyhydroxyethylene-4-laurylether (Brij 30), polyhydroxyethylene-23-laurylether (Brij 35), polyhydroxyethylene-8-stearate (Myrj 45, Cremophor AP),. . . .

DETD

120 mg phosphatidylcholine from soy-bean
 (purity > 95%)
 20 mg sodium-cholate p.a. (L/D = 3.2)
 150 µl **ethanol** (96%)
 1.45 ml Actrapid HM 100 (recombinant human insulin
 100 I.U./ml)

DETD

216 mg phosphatidylcholine from soy-bean (487 µl of a
 50% solution in absolute **ethanol**)
 27 mg phosphatidylglycerol from egg (98%)
 29.45 mg oleic acid, puriss.

3 ml Actrapid HM 100 (recombinant human insulin 100
 I.U. /ml)
40 µl 1N NaOH
20 µl 1N NaCl

DETD The lipids are dissolved in a glass vial in 0.15 ml abs. **ethanol**
 and then combined with a standard insulin solution. Further procedure is
 as described in example 239.

DETD
144.9; 152 mg phosphatidylcholine from soy-bean
24.8; 17.6 mg desoxycholate, Na-salt
1.45; 1.55 ml Actrapid HM 100 (145 I.U.)
0.16 ml **ethanol**, absolute

DETD Lipids are weighed into glass vials, dissolved with **ethanol**
 and mixed with an insulin solution. The resulting opaque suspension is
 aged over night and subsequently filtered through a 0.22. . .

CLM What is claimed is:
 . . . hydrated castor oil, sorbitanemonolaurate, lauryl- salts,
 oleoylsulfate-salts, sodium deoxycholate, sodium glycodeoxycholate,
 sodium oleate, sodium elaidate, sodium linoleate, sodium laurage,
 nonaethylene-dodecylether, **polyethylene glycol**
 -20-sorbitane-monooleate, polyhydroxyethylene-23-laurylether,
 polyhydroxyethylene-40-stearate, a sorbitane phospholipid a monolaurate
 phospholipid and a lysophospholipid.

35. The method of claim 1, wherein the agent is selected from the group
consisting of an **adrenocorticosteroid** or its analogues, an
androgen, an antiandrogen, an anabolic steroid, an anaesthetic, an
analgesic, an antiallergic, an antiarrhythmic, an antiarterosclerotic, .

IT Adrenergic agonists
IT Adrenergic antagonists
IT Allergy inhibitors
IT Anabolic agents
IT Analgesics
IT Anesthetics
IT Antiarrhythmics
IT Antibiotics
IT Anticoagulants and Antithrombotics
IT Anticonvulsants and Antiepileptics
IT Antidepressants
IT Antidiabetics and Hypoglycemics
IT Antiemetics
IT Antihistaminics
IT Antipyretics
IT Bactericides, Disinfectants, and Antiseptics
IT Bronchodilators
IT Cardiotonics
IT Cholinergic antagonists
IT Contraceptives
IT Cytotoxic agents
IT Diuretics
IT Fungicides and Fungistats
IT Ganglionic blocking agents
IT Hemostatics
IT Hypnotics and Sedatives
IT Immunomodulators
IT Inflammation inhibitors
IT Muscle relaxants
IT Mydriatics
IT Narcotics
IT Nervous system stimulants

IT Parasiticides
 IT Pesticides
 IT Poisoning
 IT Tuberculostatics
 IT Vasoconstrictors
 IT Vasodilators
 IT Virucides and Virustats
 IT Wound healing promoters
 IT Androgens
 IT Coenzymes
 IT **Corticosteroids, biological studies**
 IT Enzymes
 IT Immunoglobulins
 (transferrins containing)
 IT **Corticosteroids, biological studies**
 (glucocorticoids, transferrins containing)

L130 ANSWER 5 OF 8 USPTAFULL on STN

AN 2000:145896 USPTAFULL
 TI Formulations and methods for reducing skin irritation
 IN Hahn, Gary S., Cardiff by the Sea, CA, United States
 Thueson, David O., Poway, CA, United States
 PA Cosmederm Technologies, La Jolla, CA, United States (U.S. corporation)
 PI US 6139850 20001031 <--
 WO 9619184 19960627
 AI US 1997-860993 19970623 (8) <--
 WO 1995-US16985 19951221
 19970623 PCT 371 date
 19970623 PCT 102(e) date
 RLI Continuation-in-part of Ser. No. US 1994-362100, filed on 22 Dec 1994,
 now patented, Pat. No. US 5716625
 DT Utility
 FS Granted
 EXNAM Primary Examiner: Page, Thurman K.; Assistant Examiner: Channavajjala,
 L.
 LREP Lyon & Lyon LLP
 CLMN Number of Claims: 113
 ECL Exemplary Claim: 1
 DRWN 25 Drawing Figure(s); 13 Drawing Page(s)
 LN.CNT 1834

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Composition and methods are provided for inhibiting skin irritation
 attributable to chemical irritants or environmental conditions, by the
 application of an anti-irritant amount of aqueous-soluble strontium
 cation.
 AN 2000:145896 USPTAFULL
 PI US 6139850 20001031 <--
 WO 9619184 19960627
 AI US 1997-860993 19970623 (8) <--
 WO 1995-US16985 19951221
 19970623 PCT 371 date
 19970623 PCT 102(e) date
 SUMM . . . drug ingredients are formulated may also produce irritation in
 sensitive people, especially in the case of drugs such as topical
 corticosteroids.
 SUMM . . . as retinoids (e.g. tretinoin, retinol and retinal), carboxylic
 acids including α -hydroxy acids (e.g. lactic acid, glycolic acid),
 β -hydroxy acids (e.g. **salicylic acid**),
 α -keto acids, acetic acid and **trichloroacetic**
 acid, 1-pyrrolidone-5-carboxylic acid, capryloyl
 salicylic acid, α -hydroxy decanoic acid,
 α -hydroxy octanoic acid, gluconolactone, methoxypropyl
 gluconamide, oxalic acid, malic acid, tartaric acid, mandelic acid,

benzylic acid, gluconic. . . .

SUMM . . . prevent the irritation caused by such products. Common exfoliants include α - and β -hydroxy carboxylic acids such as lactic acid, glycolic acid, **salicylic acid** and the like, α -keto acids such as pyruvic acid, as well as assorted compounds such as acetic acid and **trichloroacetic acid**, 1-pyrrolidone-5-carboxylic acid, capryloyl **salicylic acid**, α -hydroxy decanoic acid, α -hydroxy octanoic acid, gluconolactone, methoxypropyl gluconamide, oxalic acid, malic acid, tartaric acid, mandelic acid, benzylic acid, gluconic. . . .

DETD . . . topical application of the hydroxy acid skin irritant lactic acid as well as the skin irritants glycolic acid, capsaicin, capryloyl **salicylic acid**, benzoyl peroxide, and post-shaving-applied seawater. Formulations containing the strontium cation are useful in suppressing a wide range of topical-product-induced irritation. . . . phenols, peroxides and similar irritants found in over-the-counter topical products for home or cosmetologist use (such as, 1-pyrrolidone-5-carboxylic acid, capryloyl **salicylic acid**, α -hydroxy decanoic acid, α -hydroxy octanoic acid, gluconolactone, methoxypropyl gluconamide, oxalic acid, malic acid, tartaric acid, mandelic acid, benzylic acid, and. . . or even higher) dosage forms of such irritants. The irritation attributable to combinations of such irritating ingredients, such as lactic acid/**salicylic acid** combinations and hydroxy acid/retinoid combinations, as well as irritation attributable to purified isomeric forms of such ingredients, can also be. . . .

DETD . . . example in which the vehicle has a density of 0.93 g/ml (as in a 50:50 [by volume] mixture of 95% **ethyl alcohol** and water) and the cation component is incorporated in the form of strontium nitrate (formula weight 212), representative molarity concentration. . . .

DETD . . . components) as water; organic solvents such as alcohols (particularly lower alcohols readily capable of evaporating from the skin such as **ethanol**), glycols (such as glycerin), aliphatic alcohols (such as lanolin); mixtures of water and organic solvents (such as water and alcohol),. . . .

DETD . . . skin irritation caused by certain severe skin irritants, including particularly lactic acid and glycolic acid (which are hydroxy acids), capryloyl **salicylic acid** (a β -hydroxy acid ester) and capsaicin (an isolate from cayenne and paprika known for its skin-irritating properties). The trials were. . . .

DETD . . . of the subjects. In the majority of the tests, the irritant composition was 7.5% lactic acid dissolved in a 10% **ethanol** -in-water solution.

DETD . . . of the irritant composition. Controls were performed by applying corresponding formulation(s) (pretreatment and/or skin-irritant composition) with an equimolar amount of **sodium chloride** to a contralateral portion of the subject's skin. Typically, the test materials were applied to the face of the subject.

DETD

Time Zero Tests

			Percent	
Cation	Anion	Salt Formula	Vehicle	Inhibition
<hr/>				
Strontium				
	Chloride	SrCl.sub.2	10% EtOH	58
Strontium				
	Nitrate	Sr(NO.sub.3).sub.2	10% EtOH	64

DETD . . . test compounds of the invention were formulated in Elizabeth

Arden "Visible Difference Refining Toner", with the Toner mixed with equimolar **sodium chloride** serving as the control.

The test solutions (and control) were provided in coded vials for application to either the right. . .

DETD Following a protocol parallel to that of the lactic acid irritant trials described above, glycolic acid (6.0% in 10% **ethanol-in-water**) was applied as a skin irritant to subject panels. Strontium nitrate was co-administered as an anti-irritant (time zero testing), and. . .

DETD . . . subject females. The control solution was Vaseline Smooth Legs and Feet Lotion (containing water, lactic acid (5%), glycerin, isopropyl palmitate, **PEG-40** stearate, cetyl alcohol, potassium hydroxide, steareth-2, magnesium aluminum silicate, lecithin, soya sterol, tocopheryl acetate, tetinyl palmitate, dimethicone, **menthol**, camphor, stearic acid, laureth-7, xanthan gum, polyacrylamide, C13-14 isoparaffin, corn oil, fragrance, DMDM hydantoin, iodopropynyl butylcarmamate, disodium EDTA, PG, and. . .

DETD . . . (Dow Corning, "DC344"), 7.5 ml cyclomethicone/dimethiconol (Dow Corning, "DC 1401"), 7.5 ml cyclomethicone/dimethicone copolyol (Dow Corning, "DC3225C") and 8 ml **PEG-8** and blended for 2-3 minutes. Imidizolidinyl urea (0.5%) was added as a preservative. A clear, thick gel resulted (50 ml).

DETD . . . of the salt with Mary Kay Revival Serum (with 15% lactic acid) and L'Oreal Vichy Novactia Cream (with 2% capryloyl **salicylic acid**), respectively.

CLM What is claimed is:

14. The composition of claim 11 wherein said irritant ingredient comprises **salicylic acid** or a salt thereof.

15. The composition of claim 11 wherein said irritant ingredient comprises a combination of lactic acid and **salicylic acid**, or salts thereof.

16. The composition of claim 11 wherein said irritant ingredient comprises capryloyl **salicylic acid** or a salt thereof.

. . . composition of claim 11 wherein said irritant ingredient comprises one or more of the group consisting of 1-pyrrolidone-5-carboxylic acid, capryloyl **salicylic acid**, α -hydroxy decanoic acid, α -hydroxy octanoic acid, gluconolactone, methoxypropyl gluconamide, oxalic acid, malic acid, tartaric acid, mandelic acid, benzylic acid, gluconic. . .

22. The composition of claim 11 wherein said irritant ingredient comprises **trichloroacetic acid** or a salt thereof.

IT 50-21-5, Lactic acid, biological studies 57-13-6, Urea, biological studies 58-08-2, Caffeine, biological studies 64-19-7, Acetic acid, biological studies 68-26-8, Retinol 69-72-7, Salicylic acid, biological studies 76-03-9, Trichloroacetic acid, biological studies 76-93-7, biological studies 77-92-9, Citric acid, biological studies 79-14-1, Glycolic acid, biological studies 87-69-4, biological studies 90-64-2, Mandelic acid 90-80-2 94-36-0, Benzoyl peroxide, biological studies 97-59-6, Allantoin 98-79-3 108-95-2, Phenol, biological studies 116-31-4, Retinal 127-17-3, Pyruvic acid, biological studies 144-62-7, Ethanedioic acid, biological studies 302-79-4, Tretinoin 404-86-4, Capsaicin. 515-69-5, α -Bisabolol 526-95-4, D-Gluconic acid 617-73-2, α -Hydroxy octanoic acid 1405-86-3, Glycyrrhizic acid 5393-81-7, α -Hydroxy decanoic acid 6915-15-7, Malic acid 70424-62-3 126094-21-1
(strontium compds. for reducing skin irritation due to ingredients in compns.)

AN 1998:14485 USPATFULL
 TI Formulations and methods for reducing skin irritation
 IN Hahn, Gary Scott, Cardiff by the Sea, CA, United States
 Thueson, David Orel, Poway, CA, United States
 PA Cosmederm Technologies, La Jolla, CA, United States (U.S. corporation)
 PI US 5716625 19980210 <--
 AI US 1994-362100 19941221 (8) <--
 DT Utility
 FS Granted
 EXNAM Primary Examiner: Gardner, Salle M.
 LREP Lyon & Lyon LLP
 CLMN Number of Claims: 54
 ECL Exemplary Claim: 1
 DRWN 25 Drawing Figure(s); 13 Drawing Page(s)
 LN.CNT 1646
 CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Topical formulations comprising an anti-irritant amount of aqueous-soluble strontium (Sr.sup.2+) cation, and methods for using the same to inhibit skin irritation, are disclosed.

AN 1998:14485 USPATFULL
 PI US 5716625 19980210 <--
 AI US 1994-362100 19941221 (8) <--

SUMM . . . which the active drug ingredients are formulated may also produce irritation in sensitive people, especially in drugs such as topical **corticosteroids**.

SUMM . . . as retinoids (e.g. tretinoin, retinol and retinal), carboxylic acids including α -hydroxy acids (e.g. lactic acid, glycolic acid), β -hydroxy acids (e.g. **salicylic acid**), α -keto acids, acetic acid and **trichloroacetic acid**, 1-pyrrolidone-5-carboxylic acid, capryloyl **salicylic acid**, α -hydroxy decanoic acid, α -hydroxy octanoic acid, gluconolactone, methoxypropyl gluconamide, oxalic acid, malic acid, tartaric acid, mandelic acid, benzylic acid, gluconic. . .

SUMM . . . the irritation caused by such products. Common exfolients include α - and β -hydroxy carboxylic acids such as lactic acid, glycolic acid, **salicylic acid** and the like, α -keto acids such as pyruvic acid, as well as assorted compounds such as acetic acid and **trichloroacetic acid**, 1-pyrrolidone-5-carboxylic acid, capryloyl **salicylic acid**, α -hydroxy decanoic acid, α -hydroxy octanoic acid, gluconolactone, methoxypropyl gluconamide, oxalic acid, malic acid, tartaric acid, mandelic acid, benzylic acid, gluconic. . .

DETD . . . topical application of the hydroxy acid skin irritant lactic acid as well as the skin irritants glycolic acid, capsaicin, capryloyl **salicylic acid**, benzoyl peroxide, and post-shaving-applied seawater. Formulations containing the strontium cation are useful in suppressing a wide range of topical-product-induced irritation. . . phenols, peroxides and similar irritants found in over-the-counter topical products for home or cosmetologist use (such as, 1-pyrrolidone-5-carboxylic acid, capryloyl **salicylic acid**, α -hydroxy decanoic acid, α -hydroxy octanoic acid, gluconolactone, methoxypropyl gluconamide, oxalic acid, malic acid, tartaric acid, mandelic acid, and benzylic acid),. . . or even higher) dosage forms of such irritants. The irritation attributable to combinations of such irritating ingredients, such as lactic acid/**salicylic acid** combinations and hydroxy acid/retinoid combinations, as well as irritation attributable to purified isomeric forms of such ingredients, can also be. . .

DETD . . . example in which the vehicle has a density of 0.93 g/ml (as in a 50:50 [by volume] mixture of 95% **ethyl alcohol** and water) and the cation component is incorporated in the form of strontium nitrate (formula weight 212), representative molarity concentration. . .

DETD . . . components) as water; organic solvents such as alcohols (particularly lower alcohols readily capable of evaporating from the skin such as **ethanol**), glycols (such as glycerin), aliphatic alcohols (such as lanolin); mixtures of water and organic solvents (such as water and alcohol),. . .

DETD . . . skin irritation caused by certain severe skin irritants, including particularly lactic acid and glycolic acid (which are hydroxy acids), capryloyl **salicylic acid** (a β -hydroxy acid ester) and capsaicin (an isolate from cayenne and paprika known for its skin-irritating properties). The trials were. . .

DETD . . . of the subjects. In the majority of the tests, the irritant composition was 7.5% lactic acid dissolved in a 10% **ethanol**-in-water solution.

DETD . . . of the irritant composition. Controls were performed by applying corresponding formulation(s) (pretreatment and/or skin-irritant composition) with an equimolar amount of **sodium chloride** to a contralateral portion of the subject's skin. Typically, the test materials were applied to the face of the subject.

DETD

TIME ZERO TESTS

			Percent
Cation	Anion	Salt Formula	Vehicle Inhibition
<hr/>			
Strontium			
	Chloride	SrCl.sub.2	
		10% EtOH	
			58
Strontium			
	Nitrate	Sr(NO.sub.3).sub.2	
		10% EtOH	
			64

DETD . . . test compounds of the invention were formulated in Elizabeth Arden "Visible Difference Refining Toner", with the Toner mixed with equimolar **sodium chloride** serving as the control. The test solutions (and control) were provided in coded vials for application to either the right. . .

DETD Following a protocol parallel to that of the lactic acid irritant trials described above, glycolic acid (6.0% in 10% **ethanol**-in-water) was applied as a skin irritant to subject panels. Strontium nitrate was co-administered as an anti-irritant (time zero testing), and. . .

DETD . . . subject females. The control solution was Vaseline Smooth Legs and Feet Lotion (containing water, lactic acid (5%), glycerin, isopropyl palmitate, **PEG**-40 stearate, cetyl alcohol, potassium hydroxide, steareth-2, magnesium aluminum silicate, lecithin, soya sterol, tocopheryl acetate, tetinyl palmitate, dimethicone, **menthol**, camphor, stearic acid, laureth-7, xanthan gum, polyacrylamide, C13-14 isoparaffin, corn oil, fragrance, DMDM hydantoin, iodopropynyl butylcarmamate, disodium EDTA, PG, and. . .

DETD . . . cyclomethicone (Dow Corning, "DC344"), 7.5 ml cyclomethicone/dimethiconol (Dow Corning, "DC1401"), 7.5 ml cyclomethicone/dimethicone copolyol (Dow Corning, "DC3225C") and 8 ml **PEG**-8 and blended for 2-3 minutes. Imidizolidinyl urea (0.5%) was added as a preservative. A clear, thick gel resulted (50 ml).

DETD . . . of the salt with Mary Kay Revival Serum (with 15% lactic acid) and L'Oreal Vichy Novactia Cream (with 2% capryloyl **salicylic acid**), respectively.

CLM What is claimed is:

13. The composition of claim 1 wherein said irritant ingredient comprises **salicylic acid** or a salt thereof.

14. The composition of claim 1 wherein said irritant ingredient comprises a combination of lactic acid and **salicylic acid**, or salts thereof.

15. The composition of claim 1 wherein said irritant ingredient comprises capryloyl **salicylic acid** or a salt thereof.

composition of claim 1 wherein said irritant ingredient comprises an ingredient selected from the group consisting of 1-pyrrolidone-5-carboxylic acid, capryloyl **salicylic acid**, α -hydroxy decanoic acid, α -hydroxy octanoic acid, gluconolactone, methoxypropyl gluconamide, oxalic acid, malic acid, tartaric acid, mandelic acid, benzylic acid, gluconic.

21. The composition of claim 1 wherein said irritant ingredient comprises **trichloroacetic acid** or a salt thereof.

IT 50-21-5, Lactic acid, biological studies 57-13-6, Urea, biological studies 58-08-2, Caffeine, biological studies 64-19-7, Acetic acid, biological studies 68-26-8, Retinol 69-72-7, Salicylic acid, biological studies 76-03-9, Trichloroacetic acid, biological studies 76-93-7, biological studies 77-92-9, Citric acid, biological studies 79-14-1, Glycolic acid, biological studies 87-69-4, Tartaric acid, biological studies 90-64-2, Mandelic acid 90-80-2, Gluconolactone 94-36-0, Benzoyl peroxide, biological studies 97-59-6, Allantoin 98-79-3 108-95-2, Phenol, biological studies 116-31-4, Retinal 127-17-3, Pyruvic acid, biological studies 144-62-7, Oxalic acid, biological studies 302-79-4, Tretinoin 515-69-5, α -Bisabolol 526-95-4, Gluconic acid 543-94-2, Strontium acetate 617-73-2, α -Hydroxyoctanoic acid 1405-86-3, Glycyrrhizic acid 5393-81-7, α -Hydroxydecanoic acid 6915-15-7, Malic acid 7440-24-6, Strontium, biological studies 7759-02-6, Strontium sulfate 10042-76-9, Strontium nitrate 10476-85-4, Strontium chloride 70424-62-3 126094-21-1
(strontium cation formulations for reducing skin irritation)

L130 ANSWER 7 OF 8 USPATFULL on STN

AN 90:61429 USPATFULL

TI Delivery systems for pharmaceutical or therapeutic actives

IN Partain, III., Emmett M., Bound Brook, NJ, United States

Brode, II., George L., Bridgewater, NJ, United States

PA Union Carbide Chemicals and Plastics Company Inc., Danbury, CT, United States (U.S. corporation)

PI US 4946870 19900807 <--

AI US 1988-268871 19881108 (7) <--

RLI Continuation-in-part of Ser. No. US 1988-189312, filed on 3 Feb 1988 which is a continuation-in-part of Ser. No. US 1986-871381, filed on 6 Jun 1986, now abandoned

DT Utility

FS Granted

EXNAM Primary Examiner: Griffin, Ronald W.

LREP Gibson, Henry H.

CLMN Number of Claims: 17

ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 895

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Delivery systems containing at least one aminopolysaccharide derivative are provided for the delivery of pharmaceutical or therapeutic actives to a desired topical or mucous membrane site in a subject, and wherein upon delivery, the systems provides a biocompatible, substantive, gas permeable, film from which actives are available at the designated site.

AN 90:61429 USPATFULL

PI US 4946870 19900807 <--
 AI US 1988-268871 19881108 (7) <--
 SUMM . . . is insoluble, and in which the aminopolysaccharide derivative is insoluble. Illustrative organic compound which can be employed include acetone, methanol, **ethanol**, n-propanol, isopropanol, tertiary butyl alcohol, acetonitrile, tetrahydrofuran, dioxane, 2-ethoxyethanol, dimethoxyethane, and the like.

SUMM . . . polymer is prepared by reacting a finely ground slurry of chitosan with PCA in a polar solvent such as aqueous **ethanol**, or other suitable solvent that will dissolve PCA. As indicated in the parent applications, chitosonium pyrrolidone carboxylate has a large.

SUMM . . . method for preparing chitosan salts is applicable to other organic acids that are soluble in polar organic solvents such as **ethanol**. For example, glycolic acid in aqueous **ethanol** can be reacted with chitosan to give the glycolate salt, which is also useful as a delivery system.

SUMM Anti-inflammatory analgesics such as **salicylic acid**, salicylate esters and salts, acetylsalicylic acid, diflunisal, acetaminophen, phenylbutazone, oxyphenbutazone, sulfinpyrazone, indomethacin, sulindac, fenoprofen, flurbiprofen, ibuprofen, ketoprofen, naproxen, mefenamic acid, . . .

SUMM Anti-inflammatory **corticosteroids** such as progesterone, hydrocortisone, prednisone, fludrocortisone, **triamcinolone**, **dexamethasone**, **betamethasone**, fluocinolone, and the like.

SUMM Kerolytic agents such as benzoyl peroxide, **salicylic acid**, **trichloroacetic acid**, and piroctone, and wart treatment compounds such as salicyclic acid, **trichloroacetic acid** and lactic acid, singularly or in combination with anti-viral agents.

SUMM . . . percent of the system with the remainder being a diluent and optionally, other additives. Suitable diluents include among others, water, **ethanol**, aqueous **ethanol**, isopropanol, glycerine, dimethylether, carbon dioxide, butane, **polyethylene glycol**, ethoxylated or propoxylated glucose, sorbitol derivatives, and the like.

SUMM . . . the present invention the active or actives are dissolved or suspended in an appropriate solvent or diluent such as water, **ethyl alcohol**, isopropyl alcohol, diethylether, dimethylether, acetone, ethyl acetate, or mixtures thereof, and mixed with a solution or suspension of the desired. . . Other adjuvant ingredients such as glycerine, propylene glycol, sorbitol, preservatives, stearic acid, cetyl alcohol, other high molecular weight alcohols, surfactants, **menthol**, eucalyptus oil, other essential oils, fragrances, penetration enhancers, and the like to give stable cremes, ointments, lotions, aerosols, solutions, may. . .

SUMM In the following examples, distilled water and absolute **ethanol** were used as indicated. The active is dissolved in alcohol or alcohol/water, and mixed with an aqueous solution of the. . .

DETD . . . was prepared and mixed with chitosan in varying ratios. 3.36 g of PCA was dissolved in 75 ml of absolute **ethanol**. Three 125-ml Erlenmeyer flasks were charged with 2.5 g of 0.5 mm mesh chitosan having a degree of deacetylation of. . . ml of the alcoholic PCA solution were added to each, respectively, and the slurry diluted to 50 ml with absolute **ethanol**. Each slurry was stirred for 2 hours. The molar ratios of the three solutions were respectively 1:1, 0.67:1, and 0.33:1. . .

DETD The three 2.5 g recovered chitosan samples were combined and placed in a 250-ml beaker with 100 ml of 95% **ethanol** (7.5 g in 100 ml).

DETD 3.5 g of PCA were dissolved in 16 ml of water, and the acid solution was added to the **ethanol** slurry of chitosan (7.5 g). The chitosan became swollen and curd-like. The slurry was stirred for a few minutes,

and 80 ml of 95% **ethanol** were added. The curd-like polymer precipitated, and the slurry was vacuum-filtered. By the consistency of the polymer, recovery could be. . .

DETD . . . (degree of deacylation about 0.80, ground to 0.5 mm) and 25 ml of isopropanol. A solution of 2.45 g of **salicylic acid** and 24 ml of isopropanol was added to the slurry, followed by 15 ml of water. The slurry began to. . .

DETD 0.20 g of minoxidil and 0.12 g of nicotinic acid are dissolved in a solution of 3.0 g of **ethanol** and 1.7 g of water. 5.0 g of 2.0% chitosonium niacinate in 90:10 water/**ethanol** are added, and after vigorous mixing, a clear colorless solution was obtained which is useful as a scalp/hair lotion to. . .

DETD 0.15 g of ethyl 4-aminobenzoate (benzocaine) are dissolved in 3.85 g of **ethyl alcohol** and 1.0g of water. 5.0 g of 2.0% aqueous chitosonium pyrrolidone carboxylate are added, and after vigorous mixing, a clear,. . .

DETD 0.055 g of chloamphenicol are dissolved in 2.0 g of **ethyl alcohol** and 2.95 g of water. 5.0 g of 2.0% aqueous chitosonium pyrrolidone carboxylate are added, and after vigorous mixing, a. . .

DETD 0.027 g of sulfadiazine are dissolved in 4.73 g of **ethyl alcohol**, and mixed with 5.0 g of 2% aqueous chitosonium pyrrolidone carboxylate, giving a clear colorless solution (0.27% sulfadiazine). Sulfadiazine is. . .

DETD 0.06 g of miconazole nitrate are dissolved in 4.5 g of **ethyl alcohol** and 0.44 g of water, and mixed with 5.0 g of 2% chitosonium pyrrolidone carboxylate in 90:10 water/alcohol, giving a. . .

DETD Preparation of chitosan-based **corticosteroid** lotion

DETD 0.013 g of hydrocortisone are dissolved in 4.99 g of **ethyl alcohol**, and mixed with 5.0 g of 2% aqueous chitosonium pyrrolidone carboxylate, giving a clear, colorless solution. This solution (0.13% hydrocortisone) is useful in the topical treatment of a variety of local inflammatory diseases and pruritis. Substituting 0.015 g of **dexamethasone** for 0.013 g of hydrocortisone in this formulation yields a clear, colorless solution of 0.15% **dexamethasone**, a fluorinated steroid, also used in the treatment of topical inflammatory diseases and general inflammation.

DETD 0.50 g of ibuprofen are dissolved in 4.5 g of **ethyl alcohol**, and mixed with 5.0 g of 2% chitosonium niacinate in 90:10 water/alcohol, giving a clear, colorless solution (5.0% ibuprofen). This. . .

DETD 0.01 g of retinoic acid are dissolved in 4.99 g of **ethyl alcohol**, and mixed with 5.0 g of 2% aqueous chitosonium pyrrolidone carboxylate, and vigorously shaken. With trans-retinoic acid (0.1% retinoic acid). . .

DETD Preparation of chitosan-based topical antioxidant 0.02 g of alpha-tocopherol are dissolved in 6.0 g of **ethyl alcohol**, and mixed with 4.0 g of 2% aqueous chitosonium salicylate to give a translucent, opalescent, homogeneous white fluid. This lotion. . .

DETD 2.0 g of **salicylic acid** are dissolved in 5.0 g of **ethyl alcohol** and mixed with 3.0 g of 10% aqueous chitosonium salicylate (very low molecular weight, 1% solution=5cP at 20° C.) to give a clear, colorless solution of 20% **salicylic acid**. This solution is useful as a kerolytic lotion for the treatment of acne, psoriasis, and similar skin diseases.

DETD 0.05 g of erthromycin and 0.009 g of 2-pyrrolidone-5-carboxylic acid are dissolved in 5.00 g of absolute **ethanol**. 5.00 g of 2.0% aqueous chitosonium pyrrolidone carboxylate are added, and after vigorous mixing, a clear, colorless solution is obtained,. . .

DETD Preparation of a chitosan-based **corticosteroid** lotion

DETD 0.0225 g of **triamcinolone** acetonide are dissolved in 4.99 g of **ethyl alcohol** and mixed with 5.09 g of 2% aqueous

chitosonium lactate, giving a clear, colorless solution. This solution (0.225% triamcinolone acetonide) is useful in the topical treatment of a variety of local inflammatory diseases.

IT 50-21-5DP, DL-Lactic acid, aminopolysaccharide salts 50-78-2DP, aminopolysaccharide salts 50-81-7DP, L-Ascorbic acid, aminopolysaccharide salts 56-84-8DP, L-Aspartic acid, aminopolysaccharide salts 56-86-0DP, Glutamic acid, aminopolysaccharide salts 59-67-6DP, Nicotinic acid, aminopolysaccharide salts 64-18-6DP, Formic acid, aminopolysaccharide salts 64-19-7DP, Acetic acid, aminopolysaccharide salts 68-11-1DP, Thioglycolic acid, aminopolysaccharide salts 69-72-7DP, aminopolysaccharide salts 74-87-3DP, Methyl chloride, reaction products with chitosan 75-00-3DP, Ethyl chloride, reaction products with chitosan 75-21-8DP, Oxirane, reaction products with chitosan 75-56-9DP, reaction products with chitosan 79-11-8DP, Chloroacetic acid, reaction products with chitosan 79-14-1DP, Glycolic acid, aminopolysaccharide salts 97-65-4DP, aminopolysaccharide salts 98-79-3DP, 2-Pyrrolidone-5-carboxylic acid, aminopolysaccharide salts 110-15-6DP, Butanedioic acid, aminopolysaccharide salts 110-16-7DP, Maleic acid, aminopolysaccharide salts 110-17-8DP, 2-Butenedioic acid (E)-, aminopolysaccharide salts 110-94-1DP, Glutaric acid, aminopolysaccharide salts 123-45-5DP, Sulfonyldiacetic acid, aminopolysaccharide salts 123-93-3DP, Thiodiacetic acid, aminopolysaccharide salts 127-17-3DP, aminopolysaccharide salts 141-82-2DP, Malonic acid, aminopolysaccharide salts 142-73-4DP, Iminodiacetic acid, aminopolysaccharide salts 543-24-8DP, N-Acetyl glycine, aminopolysaccharide salts 638-32-4DP, Succinamic acid, aminopolysaccharide salts 6915-15-7DP, DL-Malic acid, aminopolysaccharide salts 50744-78-0DP, reaction products with chitosan (preparation of, for pharmaceutical, cosmetic, and fluid separation membrane uses)

L130 ANSWER 8 OF 8 USPATFULL on STN

AN 89:82643 USPATFULL

TI 2-Trichloroacetoxy-3,4,5,6-tetrachlorobenzoic acid and compositions containing same for treating benign mammalian neoformations

IN Fedeli, Gianfranco, Milan, Italy

Diamantini, Giuseppe, Fano, Italy

Djaczenko, Wiktor, Rome, Italy

Strumillo, Maria, Rome, Italy

PA Djaczenko, Wiktor, Rome, Italy (non-U.S. individual)

Strumillo, Maria, Rome, Italy (non-U.S. individual)

PI US 4871769 19891003

AI US 1987-11468 19870205 (7)

RLI Continuation-in-part of Ser. No. US 1985-815092, filed on 9 Dec 1985, now abandoned

PRAI IT 1984-48046 19840416

DT Utility

FS Granted

EXNAM Primary Examiner: Moyer, Donald B.; Assistant Examiner: Parker, Julie K.

LREP Beveridge, DeGrandi & Weilacher

CLMN Number of Claims: 10

ECL Exemplary Claim: 1,2

DRWN 4 Drawing Figure(s); 3 Drawing Page(s)

LN.CNT 382

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Tetrachlorobenzoic acid derivative having the formula: ##STR1## (2-trichloroacetoxy-3,4,5,6-tetrachlorobenzoic acid), chemotherapeutically active against cutaneous and subcutaneous benign neoformations, process for its preparations and compositions containing the same.

IN Djaczenko, Wiktor, Rome, Italy

IN Strumillo, Maria, Rome, Italy

=> d his

(FILE 'HOME' ENTERED AT 07:33:28 ON 24 MAR 2005)
SET COST OFF

FILE 'HCAPLUS' ENTERED AT 07:34:02 ON 24 MAR 2005

L1 1 S (WO2000-IT309 OR IT99-RM465)/AP,PRN
E FAVA D/AU
E DJACZENKO/AU
L2 38 S E4,E5
E STRUMILLO/AU
L3 3 S E8,E12
E FAVA D/AU
L4 5 S E3-E5
L5 14 S 5 METHYL 2 1 METHYLETHYL CYCLOHEXANOL
L6 2 S L5 AND 1 ALPHA 2 BETA 5 ALPHA

FILE 'REGISTRY' ENTERED AT 07:40:50 ON 24 MAR 2005

L7 4 S 2216-51-5 OR 2623-23-6 OR 89-78-1 OR 1490-04-6
L8 435 S C10H200/MF AND 46.150.1/RID AND 1/NR
L9 213 S L8 AND CYCLOHEXANOL
L10 29 S L9 AND 5 METHYL
L11 17 S L10 AND 2 1 METHYLETHYL
L12 15 S L11 NOT LABELED
L13 16 S L7,L12
L14 14 S L10 NOT L13
L15 15 S L13 NOT ACETATE
L16 3 S L15 AND 1 ALPHA AND 2 BETA AND 5 ALPHA
L17 12 S L15 NOT L16

FILE 'HCAPLUS' ENTERED AT 07:48:08 ON 24 MAR 2005

L18 5879 S L16
L19 10732 S MENTHOL
L20 12031 S L5,L6,L18,L19

FILE 'REGISTRY' ENTERED AT 07:50:33 ON 24 MAR 2005

L21 1 S 76-03-9
L22 1 S 69-72-7

FILE 'HCAPLUS' ENTERED AT 07:51:11 ON 24 MAR 2005

L23 7287 S L21
L24 12797 S (TRICHLOROACETIC OR TRICHLORO ACETIC OR TRI CHLOROACETIC OR T
L25 29 S L20 AND L23,L24
L26 24150 S L22
L27 60605 S 2() (HYDROXYBENZOIC OR HYDROXY BENZOIC) ()ACID OR 2() (HYDROXYBE
L28 523 S L20 AND L26,L27
L29 7 S L25 AND L28

FILE 'REGISTRY' ENTERED AT 07:53:54 ON 24 MAR 2005

L30 2 S MENTHOL/CN

FILE 'HCAPLUS' ENTERED AT 07:54:06 ON 24 MAR 2005

L31 6340 S L30 OR L17
L32 19 S L31 AND L23,L24
L33 325 S L31 AND L26,L27
L34 8 S L28,L33 AND L25,L32
L35 8 S L29,L34

FILE 'REGISTRY' ENTERED AT 07:54:44 ON 24 MAR 2005

L36 1 S 25322-68-3
L37 1 S 9002-89-5
L38 1 S 112-60-7

L39 1 S 2615-15-8
L40 4 S 124-94-7 OR 378-44-9 OR 83-43-2 OR 50-02-2
L41 1 S 64-17-5
L42 1 S 7647-14-5

FILE 'HCAPLUS' ENTERED AT 07:56:05 ON 24 MAR 2005

L43 7 S L35 AND L36-L42
L44 1 S L35 AND (PVA OR POLYVINYLALCOHOL OR POLYVINYL ALCOHOL OR POLY
L45 1 S L35 AND (PEG OR POLYETHYLENEGLYCOL OR POLYETHYLENEOXIDE OR PO
L46 2 S L35 AND (TETRAETHYLENEGLYCOL OR TETRAETHYLENE GLYCOL OR TETR
L47 1 S L35 AND (HEXAETHYLENEGLYCOL OR HEXAETHYLENE GLYCOL OR HEXA ET
L48 2 S L35 AND ?CORTICOSTER?
L49 1 S L35 AND (TRIAMCINOLON? OR BETAMETHASON? OR BETA METHASON? OR
L50 5 S L35 AND (ETOH OR ETHANOL OR ETHYLALCOHOL OR ETHYL ALCOHOL)
L51 3 S L35 AND (NACL OR (NA OR SODIUM) () CHLORIDE)
L52 3 S L35,L43-L51 AND PHARMACEUT?/SC,SX,CW,BI
L53 1 S L2-L4 AND L20,L31
L54 3 S L1,L52,L53

FILE 'REGISTRY' ENTERED AT 08:01:55 ON 24 MAR 2005

L55 15 S L15,L17,L30
SEL RN
L56 355 S E1-E15/CRN
L57 0 S L56 AND 76-03-9/CRN
L58 4 S L56 AND 69-72-7/CRN

FILE 'HCAPLUS' ENTERED AT 08:03:57 ON 24 MAR 2005

FILE 'REGISTRY' ENTERED AT 08:04:22 ON 24 MAR 2005

L59 4 S L16,L30
L60 11 S L17 NOT L59
L61 4 S L59 AND MENTHOL

FILE 'WPIX' ENTERED AT 08:05:26 ON 24 MAR 2005

L62 3 S L5/BIX
L63 3577 S L19/BIX
E MENTHOL/DCN
E E3+ALL
L64 1258 S E2 OR 0557/DRN
L65 298 S E4
L66 1 S L62 AND L64,L65
E R14648+ALL/DCN
E R07025+ALL/DCN
L67 3992 S L62-L66
L68 2 S L67 AND L24/BIX
L69 1 S L67 AND (R00395/DCN OR 0395/DRN)
E TRICHLOROACETIC ACID/DCN
E E3+ALL
L70 0 S L67 AND E4
L71 236 S L67 AND L27/BIX
L72 158 S L67 AND ((R00291 OR R07025)/DCN OR 0291/DRN)
E SALICYLIC ACID/DCN
E E3+ALL
L73 0 S L67 AND E4
L74 0 S L67 AND E6
L75 3 S L67 AND E8
L76 1 S L67 AND E10
L77 0 S L67 AND E12
L78 0 S L67 AND E14
L79 0 S L67 AND E16
L80 0 S L67 AND E18
L81 1 S L67 AND E20
L82 0 S L67 AND E22

L83 0 S L67 AND E24
L84 0 S L67 AND E26
L85 8 S L67 AND E30
L86 0 S L67 AND E34
L87 1 S L67 AND E36
L88 5 S L67 AND E38
L89 1 S L68,L69 AND L71-L88
L90 181 S L67 AND (R02044/DCN OR 2044/DRN)
L91 35 S L67 AND (R01842/DCN OR 1842/DRN)
L92 2 S L67 AND (R00952/DCN OR 0952/DRN)
L93 59 S L67 AND RA01EA/DCN
L94 37 S L67 AND (R01071 OR R01242 OR R00496 OR R00002 OR R14648 OR R0
L95 45 S L67 AND (1071 OR 1242 OR 0496 OR 0002 OR 1071)/DRN
L96 236 S L67 AND (R00245 OR R01706)/DCN
L97 314 S L67 AND (0245 OR 1706)/DRN
L98 1 S L90-L97 AND L68,L69
L99 71 S L90-L97 AND L71,L72
L100 4 S L90-L97 AND L73-L87
L101 1 S L98 AND L99,L100

FILE 'WPIX' ENTERED AT 08:27:15 ON 24 MAR 2005

FILE 'USPATFULL' ENTERED AT 08:27:25 ON 24 MAR 2005

L102 1127 S L61 OR L60
L103 8487 S L5 OR L19
L104 8557 S L102,L103
L105 213 S L104 AND (L21 OR L24)
L106 2302 S L104 AND (L22 OR L27)
L107 84 S L105 AND L106
L108 5 S L107 AND L40
L109 27 S L107 AND (TRIAMCINOLON? OR BETAMETHASON? OR BETA METHASON? OR
L110 30 S L107 AND ?CORTICOSTER?
L111 7 S L107 AND CORTICOSTER?/CT
L112 33 S L108-L111
L113 0 S L112 AND L36,L37,L38,L39
L114 18 S L112 AND (PVA OR POLYVINYLALCOHOL OR POLYVINYL ALCOHOL OR POL
L115 0 S L112 AND (TETRAETHYLENEGLYCOL OR TETRAETHYLENE GLCYOL OR TETR
L116 0 S L112 AND (HEXAETHYLENEGLYCOL OR HEXAETHYLENE GLCYOL OR HEXA E
L117 28 S L112 AND (PEG OR POLYETHYLENEGLYCOL OR POLYETHYLENEOXIDE OR P
L118 32 S L114,L117
L119 33 S L112,L118
L120 33 S L119 AND (L41 OR ETOH OR ETHANOL OR ETHYLALCOHOL OR ETHYL ALC
L121 28 S L119 AND (L42 OR NACL OR (NA OR SODIUM) ()CHLORIDE)
L122 33 S L120,L121
L123 17 S L122 AND (PY<=2000 OR PRY<=2000 OR AY<=2000)
SEL AN 7 12 13 16 17
L124 5 S E1-E5 AND L123
L125 0 S L104 AND (FAVA ? OR DJACZENKO ? OR STRUMILLO ?)/AU
L126 1 S (DJACZENKO ? OR STRUMILLO ?)/AU
L127 6 S L124,L126
L128 16 S L122 NOT L123
SEL AN 10 15
L129 2 S L128 AND E6-E7
L130 8 S L127,L129 AND L102-L129

FILE 'USPATFULL' ENTERED AT 08:42:19 ON 24 MAR 2005

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